

# STIC Search Report Biotech-Chem Library

# STIC Database Tracking Number: 145345

TO: Janet Epps-Ford

Location: REM-2C05/2C18

Art Unit: 1635

Wednesday, February 16, 2005 Case Serial Number: 10/001863 From: Paul Schulwitz

**Location: Biotech-Chem Library** 

**REM-1A65** 

Phone: (571)272-2527

paul.schulwitz@uspto.gov

# **Search Notes**

Examiner Epps-Ford,

See attached results.

If you have any questions about this search feel free to contact me at any time.

Thank you for using STIC search services!

Paul Schulwitz
Technical Information Specialist
STIC Biotech/Chem Library
(571)272-2527



THIS PAGE BLANK (USPIC,

## Schreiber, David

From: Epps-Ford, Janet

Sent: Friday, February 04, 2005 1:53 PM

To: Schreiber, David

Subject: 10/001863-Score over length.

Please perform a score over length search of SEQ ID NO: 3 of application no. 10/001863.

Search for compounds that are 8 to 80 nucleobases in length, having at least 75% identity to SEQ ID NO: 3.

Please search all commerical and patent (issued and published) nucleic acid databases.

### Thanks,

Thanks,

Janet L. Epps-Ford, Ph.D.

Art Unit 1635

Mailbox: Remsen 2C18

Office: Remsen 2C05

Phone: 571-272-0757 Fax:

571-273-0757



ACCESSION: AXO57494 ACCESSION: AXO57495 ACCESSION: AXO57496 ACCESSION: AXO57499 ACCESSION: AXO57513 ACCESSION: AXO57513 ACCESSION: AXO57513	ACCESSION: AXOS7514 ACCESSION: AXOS7515 ACCESSION: AXOS7516 ACCESSION: AXOS7516 ACCESSION: AXOS7519 ACCESSION: AXOS7520 ACCESSION: AXOS7520 ACCESSION: AXOS7520 ACCESSION: AXOS7522 ACCESSION: AXOS7522 ACCESSION: AXOS7522	ACCESSION: AXAGO 25 2 2 2 ACCESSION: AXAGO 25 2 ACCESSION: BD102553 ACCESSION: AXAGO 24 8 2 ACCESSION: AXAGO 24 8 2 ACCESSION: AXAGO 24 2 ACCESSION: ARAGO 25 4 ACCESSION: ARAGO 25 4 ACCESSION: ARAGO 25 2 3 ACCESSION: ARAGO	ACCESSION: AR23793 ACCESSION: AR237134 ACCESSION: AR237134 ACCESSION: BD23134134 ACCESSION: R23929 ACCESSION: R23929 ACCESSION: AR367361 ACCESSION: AR367361 ACCESSION: AR367361 ACCESSION: AR26556 ACCESSION: AR26650 ACCESSION: AR2795182 ACCESSION: AR129530 ACCESSION: AR129530 ACCESSION: AR129530 ACCESSION: AR129530 ACCESSION: AR129533 ACCESSION: AR129533 ACCESSION: AR129533 ACCESSION: AR129533 ACCESSION: AR129533 ACCESSION: AR13128 ACCESSION: AR3333	ACCESSION: ARO46255 ACCESSION: 153307 ACCESSION: AX729602
ааааааааа	анананана	чаненнаннанн	20 1 AR32593 21 1 AR37361 21 1 AR097024 21 1 BD231344 21 1 BD231344 21 1 BC331344 21 1 BC331343 21 1 AR357361 21 1 AR357361 21 1 AR357361 20 1 AR244651 19 1 AR244651 19 1 AR244651 20 1 AR099520 20 1 AR099520 20 1 AR099520 20 1 AR123930 20 1 AR123930 20 1 AR123930 20 1 AR123930 20 1 AR123930 20 1 AR123930 20 1 AR12393 20 1 13353 20 1 13353 20 1 13353 20 1 13353 20 1 13353 20 1 AR04444 20 1 BD13692 20 1 AR04444	
888888888	288888888888888888888888888888888888888	20 20 20 19 17 17 17 16.8	669 677 777 777 777 777 777 777	15.4 15.4 15.4 15.4
	000000000			
Compugen Ltd.  Compugen Ltd.  Search time 7 Seconds  (without alignments) 3.412 Million cell undates/sec	ctcactgacaggagaacta 3	Josephan Spranner to have	Description  Total score distribution.  Description  ACCESSION.AX057533  ACCESSION.AX057533  ACCESSION.AX057533  ACCESSION.AX057531  ACCESSION.AX057531  ACCESSION.AX057531  ACCESSION.AX057507  ACCESSION.AX057507  ACCESSION.AX057483  ACCESSION.AX057483  ACCESSION.AX057486  ACCESSION.AX057486  ACCESSION.AX057486  ACCESSION.AX057486  ACCESSION.AX057486  ACCESSION.AX057486  ACCESSION.AX0574873  ACCESSION.AX0574873  ACCESSION.AX057486  ACCESSION.AX057486  ACCESSION.AX057486  ACCESSION.AX057486  ACCESSION.AX057486  ACCESSION.AX0574886  ACCESSION.AX0574886  ACCESSION.AX05748886  ACCESSION.AX0574888888888888888888888888888888888888	N: AX057 N: AX057 N: AX057 N: AX057
GenCore version pyright (c) 1993 - 2005 search, using sw model uary 16, 2005, 09:21:39	-001-863-3 gggccactgctgctcac TTY NUC 10.0 , Gapext 0.5 age, 3134 residues	80  num Match 0*  num Match 100*  num first 153 summe  ):*	SUMP SUMP SUMP SUMP SUMP 33 1 BD16116 33 1 AX05753 34 1 AX05753 24 1 AX05752 24 1 AX05752 24 1 AX05752 24 1 AX05752 23 1 AX05746 23 1 AX05746 24 1 AX05746 25 1 AX05746 27 1 AX05746 20 1 AX05746 20 1 AX05747 20 1 AX05746 20 1 AX05747 20 1 AX05747 20 1 AX05746 20 1 AX05747 20 1 AX05747 20 1 AX05747	.5 20 1
Co OM nucleic - nucleic Run on: Pebr	Title: Perfect score: 3811 Sequence: 1 acac Scoring table: IDENT Searched: 153 sc	um DB seq 16 um DB seq 16 processing: ase : Pred No. :	# dreates	0000

PI YASUNOBU YOSHIKAI, TETSI C12N15/09,C07K14/705,C12N1// C12P21/02, PC (C12N5/10,C12R1:91),C11 Description of Artificial in the control of Artificial in the cont	Query Match Best Local Similarity 100.0%; Fred Matches 33; Conservative 0; Mi Qy 114 GAAAGCTGGGAGCCCTGCGTGAGA Db 1 GAAAGCTGGGAGCCTGCGTGAGAGA	RESULT 2 AXO57533/C DEFINITION Sequence 69 from Patent WOG ACCESSION AXO57533 ACCESSION AXO57533 I GI:12310267 KEYWORDS Homo sapiens (human) ORGANISM Homo sapiens (human) ORGANISM Homo sapiens (human) ORGANISM Homo sapiens (human) ORGANISM Lome sapiens (human) ORGANISM Lome sapiens (human) ORGANISM Lome sapiens (human) AUTHORS Lorenz, E., Schwartz, D.A. and AUTHORS Variant tlr4 nucleic acid batch and Authors Variant tlr4 nucleic acid batch lowersity of Iowa Research Course of Location/Qualifies Source (homos source)  AUTHORS Variant tlr4 nucleic acid batch lowersity of Iowa Research Course of Location/Qualifies (homos source) AUTHORS Variant tlr4 nucleic acid batch lowersity of Iowa Research Course (homos second lowers) AUTHORS Variant tlr4 nucleic acid lowersity of Iowa Research Course (homos second lowers) AUTHORS Variant tlr4 nucleic acid lowersity of Iowa Research Location/Qualifies (homos lowers) AUTHORS Variant tlr4 nucleic acid lowersity of Iowa Research Location/Qualifies (homos lowers) AUTHORS Variant tlr4 nucleic acid lowersity of Iowa Research Location/Qualifies (homos lowers) AUTHORS Variant tlr4 nucleic acid lowersity of Iowa Research Location/Qualifies (homos lowers) AUTHORS VARIANT (homos lowers) AUTHORS V	Query Match Ouery Match Best Local Similarity 100.0%; Pree Matches 30; Conservative 0; M arches 30; Conservative 0; M arches 30; AccarrraAcrrcAcrrcAcrrcAcrrcAcrrcAcrrcA	SULT 3 057532 AD 057532 AD 057532 AD CESSION AD FRASION
ACCESSION: AR042291 ACCESSION: AR083517 ACCESSION: AR083519 ACCESSION: AR083519 ACCESSION: CQ788034 ACCESSION: AR150152 ACCESSION: AR150152 ACCESSION: AR150152 ACCESSION: AR150152 ACCESSION: AR150153 ACCESSION: AR333333349	ACCESSION: AR317378 ACCESSION: BD084033 ACCESSION: A86518 ACCESSION: AR030970 ACCESSION: AR108815 ACCESSION: BAL78882 ACCESSION: BD184152 ACCESSION: BD184152	ACCESSION: E13800 ACCESSION: 121071 ACCESSION: 123988 ACCESSION: 123988 ACCESSION: AR205764 ACCESSION: AR23731 ACCESSION: AR311481 ACCESSION: AR347090 ACCESSION: AX544171 ACCESSION: AX544171 ACCESSION: AX54228 ACCESSION: AX542328 ACCESSION: AX542328 ACCESSION: AX642374 ACCESSION: AX622873 ACCESSION: AX622873 ACCESSION: AX622873 ACCESSION: AX622874 ACCESSION: AR033742 ACCESSION: AR033742 ACCESSION: AR033742 ACCESSION: AX623742 ACCESSION: AX628742 ACCESSION: AX623742 ACCESSION: AX623742 ACCESSION: AX623742 ACCESSION: AX6274111		a3 bp DNA linear PAT 17-JAN-2003 echanism for preventing infection and last last last last last last last last
4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4	**************************************	2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2	0000 0000 0000 0000 0000 0000 0000 0000	BD161165  Protein participating in mechanism for pr gene encoding the protein. BD161165. BD161165. BD161165. BD 2002176986-A/3. Synthetic construct other sequences; artificial sequences. I bases 1 to 33 solution. Yoshikai, Y., Matsuguchi, T. and Iwami, K. Protein participating in mechanism for progene encoding the protein Patent: JP 2002176986-A 3 25-JUN-2002; NAGOYA INDUSTRIAL SCIENCE RESEARCH INSTIT OS Artificial Sequence NP JP 2002176986-A 3 PD 25-JUN-2002 PP 14-DEC-2000 JP 2000380561
1007 1008 1008 1100 1111 1112 1113 1114 1114 1116 1116 1116 1116	01284886	0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	4444	RESULT 1 BD161165 LOCUS DEFINITION ACCESSION VERSION KEYWORDS SOURCE ORGANISM REFERENCE AUTHORS TITLE JOURNAL COMMENT

```
PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ô
                                                                                                                                                                                                                                                                                                                           PAT 17-JAN-2001
                                                                                                                                                                                                          ö
                                    12N15/00,C12N5/00,(C12N5/00,C12R1:91) CC
. Sequence: CDNA to sequence 2 FH Key
*rs
                                                                                                                                                                                                                                                                                                                                                                                                                ata; Craniata; Vertebrata; Euteleostomi;
tes; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              and Schutte,B.C.
d and uses thereof
21-DEC-2000;
rch Foundation (US) ; Lorenz, Eva (US)
iers
                                                                                                                                                                                                                                                                                                                                                                                                                                                     and Schutte, B.C.
1 and uses thereof
11.DEC-2000;
cch Foundation (US); Lorenz, Eva (US)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
SUYA MATSUGUCHI, KENICHIRO IWAMI PC
/15,C12N1/19,C12N1/21,C12N5/10// PC
                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                .
0
                                                                                                                                                                                                             ö
                                                                                         nism='Artificial Sequence'.
                                                                                                                                                                                                                                                                                                                             linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    re 30; DB 1; Length 30;
ed. No. 8.4;
Mismatches 0; Indels
                                                                                                                                                                                re 33; DB 1; Length 33; ed. No. 5; Nismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            hetic construct"
signed DNA"
                                                                                                                                netic construct"
nic DNA"
:32630"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                DNA
                                                                                                                                                                                                                                      SACTTGGCCC 146
                                                                                                                                                                                                                                                                                                                               DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          al sequences.
                                                                                                                                                                                                                                                     TTCAGGT 1735
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     sapiens"
signed DNA"
:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                31 bp
WO0077204.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        |||||||
TTCAGGT 1
                                                                                                                                                                                                                                                                                                                             30 bp
100077204.
```

PAT 17-JAN-2001

VERSION KEYWORDS SOURCE ORGANISM

REFERENCE AUTHORS TITLE

JOURNAL

source

FEATURES

DEFINITION ACCESSION

RESULT 4 AX057530 LOCUS

a

```
PAT 27-SEP-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       PAT 17-JAN-2001
                                                                                                                                                                          Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Taira, K. and Kawasaki, H. cells
Patent: WO 200407622-A 162 10-SEP-2004;
National Institute of Advanced Industrial Science and Tec hnology
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                       Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 43 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                  0.7%; Score 26; DB 1; Length 26; ilarity 100.0%; Pred. No. 16; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Length 24;
                                                       linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ch 0.6%; Score 24; DB 1; I Similarity 100.0%; Pred. No. 22; 24; Conservative 0; Mismatches (
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           CO873743 24 bp RNA
Sequence 162 from Patent WO2004076622.
CO873743. GI:52747335
                                                       DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DNA

    .26
    /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         /mol_type="unassigned RNA" /db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 24 bp
Sequence 41 from Patent WO0077204.
AX057505
AX057505.1 GI:12310239
                                                Sequence 43 from Patent WO0077204. AX057507
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1. .24
/organism="Homo sapiens"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  2278 TCTGCCTTCACTACAGAGACTTTA 2301
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  24
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1 rcreccrrcacracagagacrrra
                                                                                                      AX057507.1 GI:12310241
                                                                                                                                        Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens (human)
Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens
                                                                                                                                                            Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
Best Local Similarity
Matches 26; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match
Best Local &
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       KEYWORDS
SOURCE
ORGANISM
                                                                  DEFINITION
ACCESSION
VERSION
KEYWORDS
                 RESULT 6
AX057507/c
LOCUS
                                                                                                                                                                                                                                                                                                                   source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 8
AX057505/c
LOCUS
                                                                                                                                        SOURCE
                                                                                                                                                                                                         REFERENCE
AUTHORS
TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 7
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             CQ873743
                                                                                                                                                                                                                                                                                               FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ઠ
                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      PAT 17-JAN-2001
                                                                                                                                                                                                                                             PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sec. 35.
                                                                                                                                                                                                                                                                                                                                                                                           Lorenz,E., Schwartz,D.A. and Schutte,B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 66 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Lorenz, B., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 67 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                     Length 31;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DB 1; Length 30;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     0.7%; Score 27; DB 1; Length 27;
100.0%; Pred. No. 14;
ive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                             linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        linear
                                                                                        Indels
                                                     DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1. .30
/organism="synthetic construct"
/mol type="unassigned DNA"
/db_xref="taxon:32630"
/note="A primer"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DNA
                                                                                                                      1329 GGTTGCTGTTCTCAAAGTGATTTTGGGACAA 1359
                                                                                                                                                                                                                                                                                                                            synthetic construct
synthetic construct
other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1030 GATTAGCATACTTAGACTACTCGATG 1059
                                                                                                                                          GGTTGCTGTTCTCAAAGTGATTTTTGGGAGAA 31
                                                    Query Match

0.8%; Score 29.4; Di
Best Local Similarity 96.8%; Pred. No. 10;
Matches 30; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Score 28.4; D
Pred. No. 12;
0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1. .27
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1 GATTAGCATACTTAGACTACTACCTCCATG 30
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    27 bp
Sequence 67 from Patent WO0077204.
AX057531
                                                                                                                                                                                                                                       30 bp
Sequence 66 from Patent WO0077204.
AX057530
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1252 GIGGGAATGCTTTTCAGAAGTTGATC 1278
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               27 GTGGGAATGCTTTTTCAGAAGTTGATC 1
/db_xref="taxon:32630"
/note="A_primer"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       AX057531.1 GI:12310265
                                                                                                                                                                                                                                                                                             AX057530.1 GI:12310264
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         0.7%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
Best Local Similarity 96./v
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Best Local Similarity 100.
Matches 27; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
```

RESULT 5 AX057531/c LOCUS

ઠ

REFERENCE AUTHORS

JOURNAL

TITLE

FEATURES

ઠે

ö

Gaps

ö

ö

Gaps

.; 0

셤

```
PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       PAT 17-JAN-2001
PR 31-MAR-2000 JP 00P 99617,22-NOV-2000 JP 00P 356719 PR 28-MAR-2001 US 09/806158
FI SHOJI FURUSAKO, SADAO MORI, KAMON SHIRAKAWA, TOMOHIRO TAKAHASHI PC C12N15/00, C07K7/08, C07K14/705, C07K16/28, A61K45/00, A61P31/04, PC A61P39/395, A61P43/00, G01N33/15, G01N33/50, G01N33/577 CC TLR/CD14 binding inhibitor FH Key Location/Qualifiers
FT SOURCE 1.24
/organism='Artificial Sequence'.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens
Bukaryota; Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi;
Mammalia; Butheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Lorenz, E., Schwartz, D.A. and Schutte, B.C. Variant tlr4 nucleic acid and uses thereof Patent: WO 0077204-A 23 21-DEC-2000; University of Iowa Research Foundation (US); Lorenz, Eva (US) Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 19 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          linear
                                                                                                                                                                                                                                                                                                                      ch 0.6%; Score 24; DB 1; Length 24; Similarity 100.0%; Pred. No. 22; 24; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Length 23;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Indels
                                                                                                                                                                                                                                 1. .24
/organism="synthetic construct"
/mol type="genomic DNA"
/db_xref="taxon:32630"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         0.6%; Score 23; DB 1;
100.0%; Pred. No. 26;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             100.0%; Pred. No. 26, tive 0; Mismatches

    .23
    /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    23 bp
Sequence 23 from Patent WO0077204.
AX057487
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AX057483 23 bp
Sequence 19 from Patent W00077204.
AX057483.1 GI:12310217
                                                                                                                                                                                                                                                                                                                                                                                                         1137 GACTITICITATAATITICGGAIGG 1160
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     943 TGGGAGAATTTAGAAATGAAGGA 965
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1 TGGGAGAATTTAGAAATGAAGGA 23
                                                                                                                                                                                                                                                                                                                                                                                                                              GACTTTTCTTATAATTTCGGATGG 1
                                                                                                                                                                                                        Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AX057487.1 GI:12310221
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match 0.6
Best Local Similarity 100.
Matches 23; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens
                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                                                                                                                                                                                                                                                                     Best Local
Matches 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        REFERENCE
AUTHORS
TITLE
JOURNAL
                                                                                                                                                                                                                                     source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 12
AX057487
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 11
AX057483
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               FEATURES
                                                                                                                                                                                                                 FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ଚ
                                                                                                                                                                                                                                                                                                                                                                                                                   8
                                                                                                                                                                                                                                                                                                                                                                                                                                                       d
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PAT 27-AUG-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                BD102554

N BD102554

N BD102554.

N BD102554.1 GI:22648128

WO 0172993-A/7.

Synthetic construct
Synthetic Synthetic Synthetic
Synthetic Synthetic Synthetic
Synthetic Conty, Shoil
FURVORANO, TOWOHIRO TAKAHASHI
SHIRAKAWA, TOWOHIRO TAKAHASHI
SON Artificial Sequence
NO 0172993-A/7

PD 04-OCT-2001
PP 02-APR-2001 WO 2001JP002869
                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Eukaryoča; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                    Bukaryota; Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi;
Mammalia; Butheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Lorenz, E., Schwartz, D.A. and Schutte, B.C. variant tlr4 nucleic acid and uses thereof Patent: WO 0077204-A 46 21-DEC-2000; University of Iowa Research Foundation (US); Lorenz, Eva (US) Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gaps
                                                                          Lorenz, E., Schwartz, D.A. and Schutte, B.C. Variant tlr4 nucleic acid and uses thereof Patent: WO 0077204-A 41 21-DEC-2000; University of Iowa Research Foundation (US); Lorenz, Eva (US) Location/Qualifiers
                                                                                                                                                                                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                      ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Score 24; DB 1; Length 24;
Pred. No. 22;
                                                                                                                                                                                                                                                                             ch 0.6%; Score 24; DB 1; Length 24; 1 Similarity 100.0%; Pred. No. 22; 24; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match 0.6%; Score 24; DB Best Local Similarity 100.0%; Pred. No. 22; Matches 24; Conservative 0; Mismatches

    .24
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

                                                                                                                                                                                                  organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AX057510 24 bp
Sequence 46 from Patent W00077204.
AX057510 GI:12310244
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1464 TTGAAACAAATGAGTGAGTTTTCA 1487
                                                                                                                                                                                                                                                                                                                                                                836 AGAAATTAGGCTTCATAAGCTGAC 859
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 24 rrgaaacaaargagrgagrrrrca 1
                                                                                                                                                                                                                                                                                                                                                                                    24 AGAAATTAGGCTTCATAAGCTGAC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Homo sapiens
                                                                                                                                                                                                                                                                                       Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                             AX057510/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 10
BD102554/c
                                                                                                                                                                                            source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AUTHORS
                                                                                                                                                                                                                                                                                                                                 Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AUTHORS
                                                                      REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
                                                                                                                              JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  COMMENT
                                                                                                                                                                     FEATURES
```

ö

ò 셤 S

```
PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   PAT 20-APR-2002
                                                                                               PAT 17-JAN-2001
                                                                                                                                                                                                                                       Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                               Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tir4 nucleic acid and uses thereof
Patent: WO 0077204-A 22 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204.A 42 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                               linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match 0.6%; Score 22; DB 1; Length 22; Best Local Similarity 100.0%; Pred. No. 30; Matches 22; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Length 22
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match 0.6%; Score 22; DB 1;
Best Local Similarity 100.0%; Pred. No. 30;
Matches 22; Conservative 0; Mismatches
                                                                                               DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     27 bp D1
Sequence 333 from patent US 6346398.

    .22
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

    .22
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   22 bp
Sequence 42 from Patent WO0077204.
AX057506
                                                                                               AX057486 22 bp Sequence 22 from Patent WO0077204.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1323 TTCAAAGGTTGCTGTTCTCAAA 1344
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                992 TCTAGAGGGCCTGTGCAATTTG 1013
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1 TICAAAGGITGCIGITCTCAAA 22
23 CCCACATTGAAACTCAAATCTCT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   22 TCTAGAGGCCTGTGCAATTTG 1
                                                                                                                                     AX057486
AX057486.1 GI:12310220
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AX057506.1 GI:12310240
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AR184845.1 GI:20230810
                                                                                                                                                                                                 Homo sapiens (human)
Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AR184845
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                              DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                         source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 16
AX057506/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   source
                                                                                                                                                                                                                    ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DEFINITION
                                                                                                                                                                                                                                                                          REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 17
AR184845
LOCUS
                                                        RESULT 15
AX057486
LOCUS
                                                                                                                                                                                                                                                                                                                                           JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                  FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    VERSION
윱
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    g
                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                    PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                             Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Buteleostomi,
Mammalia, Butheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Eukaryota; Metazoa, Chordata, Craniata, Vertebrata, Buteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 32 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 44 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match 0.6%; Score 23; DB 1; Length 23; Best Local Similarity 100.0%; Pred. No. 26; Matches 23; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     0.6%; Score 23; DB 1; Length 23;
100.0%; Pred. No. 26;
tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                      linear
                                                                                               Length 23
                                                                                                                                     0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  linear
                                                                                             DB 1;
. 26;
                                                                                                                                                                                                                                                                                                                      DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DNA
                                                                                           Query Match 0.6%; Score 23; DB Best Local Similarity 100.0%; Pred. No. 26; Matches 23; Conservative 0; Mismatches
                /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

    .23
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

    .23
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          23 bp
Sequence 44 from Patent WO0077204.
AX057508.1 GI:12310242
                                                                                                                                                                                                                                                                                                                    23 bp
Sequence 32 from Patent WO0077204.
                                                                                                                                                                          1410 TCAAACTTCTTGGGCTTAGAACA 1432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1200 CCCACATTGAAACTCAAATCTCT 1222
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           2459 TATCATCTTCATTGTCCTGCAGA 2481
                                                                                                                                                                                                 TCAAACTTCTTGGGCTTAGAACA 23
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1 TATCATCTTCATTGTCCTGCAGA 23
                                                                                                                                                                                                                                                                                                                                                                              AX057496.1 GI:12310230
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                      Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 23; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                           Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Best Local Similarity
Matches 23; Conserv
                                                                                                                                                                                                                                                                                                                                                           AX057496
```

Query Match

δ

RESULT 14 AX057508/c

LOCUS

ACCESSION VERSION

KEYWORDS SOURCE ORGANISM

REFERENCE AUTHORS

JOURNAL

FEATURES

AX057496 LOCUS DEFINITION

RESULT 13

a ઠે

ORGANISM ACCESSION VERSION KEYWORDS SOURCE

TITLE JOURNAL

PEATURES

AUTHORS REFERENCE

ö

Gaps

..

ö

Gaps ö

Gaps

ORGANISM

REFERENCE AUTHORS

TITLE

source

JOURNAL

FEATURES

```
PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     PAT 17-JAN-2001
                                                                                                                                                                                            PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                          Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 59 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                           Lorenz,E., Schwartz,D.A. and Schutte,B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 39 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                           ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match

0.6%; Score 21; DB 1; Length 21;
Best Local Similarity 100.0%; Pred. No. 34;
Matches 21; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.6%; Score 21; DB 1; Length 21; llarity 100.0%; Pred. No. 34; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                linear
                           Indels
                             ô
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DNA
                                                                                                                                                                                                  DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
         red. No. 34;
Mismatches

    .21
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   21 bp
Sequence 59 from Patent WO0077204.
AX057523
                                                                                                                                                                                                AX057503 21 bp
Sequence 39 from Patent WO0077204.
AX057503
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ďq
           Pred. No.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  2813 AGATATGCAGGCTGCTAATC 2833
                                                                   1093 TITCTTCATTTTCCCTGGTGA 1113
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          524 AGCCTTTTCTGGACTATCAAG 544
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1 TTCTTCATTTCCCTGGTGA 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               100.08; Pre-
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AX057523.1 GI:12310257
                                                                                                                                                                                                                                                            AX057503.1 GI:12310237
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens (human)
Homo sapiens
                                                                                                                                                                                                                                                                                              Homo sapiens (human)
             Best Local Similarity 100.
Matches 21; Conservative
                                                                                                                                                                                                                                                                                                                      Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Best Local Similarity
Matches 21; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AX057472
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 22
AX057472
LOCUS
                                                                                                                                                                  RESULT 20
AX057503/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AX057523/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                        DEFINITION
                                                                                                                                                                                                                                                                                                                      ORGANISM
                                                                                                                                                                                                                                                                                                                                                                           REFERENCE
AUTHORS
TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     FEATURES
                                                                                                                                                                                                                                                                                    KEYWORDS
                                                                                                                                                                                                                                                              VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          SOURCE
                                                                                                                                                                                                          LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ਨੇ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 셤
                                                                                                           셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                         PAT 20-APR-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Bukaryoča; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Butheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Tobase 1 to 27)
Pavco, P., McSwiggen, J., Stinchcomb, D. and Escobedo, J.
Machod and reagent for the treatment of diseases or conditions
method and reagent for the treatment of diseases or conditions
related to levels of vascular endochelial growth factor receptor
Patent: US 6346398-A 6519 12-FEB-2002;
                                   Unknows...
Unclassified.
Unclassified.
1 (bases 1 to 27)
Pavco,P., McSwiggen,J., Stinchcomb,D. and Escobedo,J.
Pavco,P., McSwiggen, for the treatment of diseases or conditions Method and reagent for the treatment of diseases or conditions related to levels of vascular endothelial growth factor receptor Patent: US 6346398-A 333 12-FEB-2002;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 20 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                        ő
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Similarity 85.2%; Score 21.2; DB 1; Length 27; Similarity 85.2%; Pred. No. 51; Conservative 0; Mismatches 4; Indels
                                                                                                                                                                                                                                            Score 21.2; DB 1; Length 27;
Pred. No. 51;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.6%; Score 21; DB 1; Length 21;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                             linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                             AR191031 27 bp DNA
Sequence 6519 from patent US 6346398.
AR191031 GI:20236996

    .21
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /organism="unknown"
/wol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          21 bp
Sequence 20 from Patent WO0077204.
AX057484
                                                                                                                                                                              1. .27
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                          0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              2673 ACATCTATCTGAAGAGGAAAATAAAA 2699
                                                                                                                                                                                                                                                                                                                                2673 ACATCTATCTGAAGAGGAAAAATAAAA 2699
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1 ACATCTGTCTGATGANGAAACATAAAA 27
                                                                                                                                                                                                                                                                                                                                                     1 ACATCTGTCTGATGANGAAATATAAAA 27
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       AX057484.1 GI:12310218
                                                                                                                                                                                                                                                        0.6%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Homo sapiens (human)
                                                                                                                                                                                                                                      Query Match
Best Local Similarity 85.4.
- المالية 23; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Unknown.
                             Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              23;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
Best Local S:
Matches 23
```

SOURCE ORGANISM

REFERENCE AUTHORS TITLE

DEFINITION

AR191031

ACCESSION VERSION KEYWORDS source

JOURNAL

FEATURES

DEFINITION

RESULT 19

AX057484 LOCUS

ACCESSION VERSION KEYWORDS SOURCE

ORGANISM

REFERENCE AUTHORS

JOURNAL

FEATURES

ö

Gaps

ö

ö

Gaps

ö

Query Match

FEATURES

```
PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
     Patent: WO 0077204-A 10 21-DEC-2000;
University of Iowa Research Foundation (US) ; Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 16 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tir4 nucleic acid and uses thereof
Patent: WO 0077204 A. 11 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                    Length 20;
                                                                                                                                                                                                                                                                                                                                                      linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ch 0.5%; Score 20; DB 1; Length 20; 1 Similarity 100.0%; Pred. No. 40; 20; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 40;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      linear
                                                                                                                                                                                        0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0; Indels
                                                                                                                                                    0.5%; Score 20; DB 1;
100.0%; Pred. No. 40;
                                                                                                                                                                                                                                                                                                                                                    DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        DNA
                                                                                                                                                        100.0%; Pred. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            100.0%; Pred. w.

    .20
    /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1. .20
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                  AX057475 20 bp
Sequence 11 from Patent WO0077204.
AX057475.1 GI:12310209
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20 bp
Sequence 16 from Patent WO0077204.
AX057480
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Location/Qualifiers
                                                                                                                                                                                                                        2467 TCATTGTCCTGCAGAAGGTG 2486
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            2590 GACGACTCAGAAAGCCCTG 2609
                                                                                                                                                                                                                                                          1 rcarrercerscasaders 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 20 GACGACTCAGAAAAGCCCTG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AX057480.1 GI:12310214
                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match
Best Local Similarity 100.
                                                                                                                                                  Query Match 0.5
Best Local Similarity 100.
Matches 20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
Best Local Similarity
Matches 20; Conserva
                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens
                                                              source
                                                                                                                                                                                                                                                                                                               RESULT 25
AX057475/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                source
                                                                                                                                                                                                                                                                                                                                                                   DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                              ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 26
AX057480
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        TITLE
         JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                      ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                       VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ACCESSION
                                           FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         VERSION
                                                                                                                                                                                                                            ઠ
                                                                                                                                                                                                                                                    g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Ş
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ద
                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Buteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                Euteleostomi;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                            Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 077204-A 8 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 9 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                             Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Butele
Mammalia, Butheria, Primates, Catarrhini, Hominidae, Homo
                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 40;
iive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      linear
                                                                                                                                                                                                                                                                                                                                                    Length 20
                                                                                                                                                                                                                                                                                                                                                                                      0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Lorenz, E., Schwartz, D.A. and Schutte, B.C. Variant tlr4 nucleic acid and uses thereof
                                                                                                                                                                                                                                                                                                                                                    DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        DNA
                                                                                                                                                                                                                                                                                                                                                 Query Match

0.5%; Score 20; DB
Best Local Similarity 100.0%; Pred. No. 40;
Matches 20; Conservative 0; Mismatches

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    20 bp
Sequence 10 from Patent WO0077204.
AX057474
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AX057473 20 bp
Sequence 9 from Patent WO0077204.
Sequence 8 from Patent WO0077204.
AX057472
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                      448 ATGGGGCATATCAGAGCCTA 467
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          587 AGAGAACTTCCCCATTGGAC 606
                                                                                                                                                                                                                                                                                                                                                                                                                                           1 ATGGGCATATCAGAGCCTA 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AGAGAACTTCCCCATTGGAC 1
                                     AX057472.1 GI:12310206
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AX057474.1 GI:12310208
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       AX057473.1 GI:12310207
                                                                           Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Conservative
                                                                                            Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
Best Local Similarity
Matches 20; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            20
                ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                            source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 24
AX057474
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AX057473/c
LOCUS
     DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ORGANISM
                                                                                                                                               REFERENCE
AUTHORS
                                                                                                                                                                                  TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 23
```

FEATURES

ઠ 셤

VERSION

a ଚ

ö

Eva (US)

ò

```
Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                Lorenz, E., Schwartz, D.A. and Schutte, B.C. Variant tlr4 nucleic acid and uses thereof Patent: WO 0077204-A 24 21-DEC-2000; University of Iowa Research Foundation (US) ; Lorenz, Location/Qualifiers

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                    1. .20
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                            0.5%; Score 20; DB
100.0%; Pred. No. 40;
ive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AX057491 20 bp
Sequence 27 from Patent WO0077204.
AX057491. GI:12310225
                                                                                                                                                                                                                                                                                                                                                                                                                              AX057489 20 bp
Sequence 25 from Patent WO0077204.
AX057489
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1612 TCCAGGAAACTTCCTTCCA 1631
                                                                                                                                                                                                                                                                                                                          1541 CAGAGITGCTTTCAATGGCA 1560
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1 TCCAGGAAACTTCCTTCCA 20
                                                                                                                                                                                                                                                                                                                                           1 CAGAGTTGCTTTCAATGGCA 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AX057489.1 GI:12310223
AX057488.1 GI:12310222
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens (human)
                             Homo sapiens (human)
Homo sapiens
                                                                                                                                                                                                                                                                                              20; Conservative
                                                                                                                                                                                                                                                                  Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          REFERENCE
AUTHORS
TITLE
JOURNAL
                                                                                                                                                                                           source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            REFERENCE
AUTHORS
TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 31
AX057491
                                                                                                                                                                                                                                                                                                                                                                                                                                                     DEFINITION
                                   SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 30
AX057489
LOCUS
                                                                                                                           TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                Matches
                                                                                            REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        FEATURES
    VERSION
KEYWORDS
                                                                                                                                                                          FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Š
                                                                                                                                                                                                                                                                                                                                ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                              PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                        Eukaryoča; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 21 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                 Lorenz, E., Schwartz, D.A. and Schutte, B.C. variant tlr4 nucleic acid and uses thereof Patent: WO 0077204-A 17 21-DEC-2000; University of Iowa Research Foundation (US); Lorenz, Eva (US) Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 40; tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                linear
                                                                                                                                                                                                                                                                                                                                                                                                                                           Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0; Indels
                                                                                                                                  linear
                                                                                                                                                                                                                                                                                                                                                                                                                                             DB 1;
40;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DNA
                                                                                                                                  DNA

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                               0.5%; Score 20; DB ilarity 100.0%; Pred. No. 40; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 20 bp
Sequence 24 from Patent WO0077204.
AX057488
                                                                                                                                                                                                                                                                                                                                                                     1. .20
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AX057485 20 bp
Sequence 21 from Patent WO0077204.
AX057485
                                                                                                                                   20 bp
Sequence 17 from Patent WO0077204.
AX057481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1190 TGGACAGTTTCCCACATTGA 1209
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           700 ACTIGGACCTTTCCAGCAAC 719
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1 TGGACAGTTTCCCACATTGA 20
                           GGTGGCTGTGGAGACAAATC 576
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1 ACTIGGACCITICCAGCAAC 20
                                                        20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AX057485.1 GI:12310219
                                                                                                                                                                                 AX057481.1 GI:12310215
                                                       gereecrereeacacaarc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens (human)
                                                                                                                                                                                                                Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
Best Local Similarity 100.
Matches 20, Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                               Best Local Similarity
Matches 20; Conserv
                                                                                                                                                                                                                                sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
                                                          г
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 DEFINITION
ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Bource
                                                                                                                                                DEFINITION
ACCESSION
VERSION
                                                                                                                                                                                                 KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                            Bource
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RESULT 29
AX057488
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                              REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                             TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 28
                                                                                                         RESULT 27
AX057481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AX057485
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          FEATURES
                                                                                                                                                                                                                                                                                                                                                          FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           용
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ò
                                                        g
```

```
.
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            PAT 18-JAN-2001
                                                                                                                                                                                                                                                           PAT 17-JAN-2001
                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens
Homo sapiens
Eukaryota, Metazoa, Chordata, Craniata, Vertebrata; Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                      Homo sapiens
Bukaryota: Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi;
Mammalia; Butheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Lorenz,E., Schwartz,D.A. and Schutte,B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 27 21-DEC-2000;
University of Iowa Research Foundation (US) ; Lorenz, Eva (US)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Lorenz,E., Schwartz,D.A. and Schutte,B.C. Variant tlr4 nucleic acid and uses thereof Patent: WO 0077204-A 25 21-DEC-2000; University of Iowa Research Foundation (US); Lorenz, Eva (US)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ..
0
                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match 0.5%; Score 20; DB 1; Length 20; Best Local Similarity 100.0%; Pred. No. 40; Matches 20; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 linear
DB 1; Length 20; 40;
                                             0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DNA
                                                                                                                                                                                                                                                                   DNA
```

PAT 17-JAN-2001

source

FEATURES

Matches

g ò

```
Homo sapiens
Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Butheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                               Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 30 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tir4 nucleic acid and uses thereof
Patent: WO 0077204-A 31 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match 0.5%; Score 20; DB 1;
Best Local Similarity 100.0%; Pred. No. 40;
Matches 20; Conservative 0; Mismatches (
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.5%; Score 20; DB 1;
100.0%; Pred. No. 40;
tive 0; Mismatches C
                                                                                                                DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DNA

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              20 bp
Sequence 33 from Patent WO0077204.
AX057497
AX057497.1 GI:12310231
                                                                                                              AX057494 20 bp
Sequence 30 from Patent WO0077204.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AX057495 20 bp
Sequence 31 from Patent WO0077204.
AX057495
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  2210 CCAGGATGAGGACTGGGTAA 2229
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2351 AAGCCGAAAGGTGATTGTTG 2370
1 CGGTCCTCAGTGTGCTTGTA 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             CCAGGATGAGGACTGGGTAA 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1 AAGCCGAAAGGTGATTGTTG 20
                                                                                                                                                                     AX057494.1 GI:12310228
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AX057495.1 GI:12310229
                                                                                                                                                                                                          Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens (human)
Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
Best Local Similarity
                                                                                                                                                                                  KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                  source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Bource
                                                                                                                              DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ORGANISM
                                                                                                                                                                                                                                                                                REFERENCE
AUTHORS
TITLE
                                                                                                                                                 ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 35
AX057495
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 36
AX057497
                                                                                                                                                                                                                                                                                                                                        JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             JOURNAL
                                                                           RESULT 34
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      VERSION
KEYWORDS
                                                                                             AX057494
                                                                                                                                                                                                                                                                                                                                                                             FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          셤
                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                    PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                   Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Lorenz, E., Schwartz, D. A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 007204-A 28 21-DEC-2000;
University of Iows Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 29 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                   Gaps
                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 40;
rative 0; Mismatches 0; Indels
                                                                                                            Length 20
                                                                                                                                                                                                                                                                                                                    linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DB 1; Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0; Indels
                                                                                                                                                 0; Indels
                                                                                                          DB 1;
                                                                                                                                                                                                                                                                                                                    DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match
Best Local Similarity 100.0%; Pred. No. 40;
Matches 20; Conservative 0; Mismatches
                                                                                                          0.5%; Score 20; DB 100.0%; Pred. No. 40; tive 0; Mismatches

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                   AX057492 20 bp
Sequence 28 from Patent WO0077204.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AX057493 20 bp
Sequence 29 from Patent WO0077204.
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Location/Qualifiers
                                                                                                                                                                                  1924 ACCAGAGTTTCCTGCAATGG 1943
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        2017 IGCCIGIGCIGAGITIGAAT 2036
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             2074 CGGTCCTCAGTGTGCTTGTA 2093
                                                                                                                                                                                                      1 TGCCTGTGCTGAGTTTGAAT 20
                                                                                                                                                                                                                                                                                                                                                      AX057492
AX057492.1 GI:12310226
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AX057493.1 GI:12310227
                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Homo sapiens (human)
Homo sapiens
                                                                                                                                               20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                 Homo sapiens
                                                                                                                              Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match
Best Local Simi
Matches 20;
                                                                                                          Query Match
```

AX057493 LOCUS DEFINITION

RESULT 33

ACCESSION VERSION

ORGANISM

KEYWORDS SOURCE

REFERENCE AUTHORS

JOURNAL

FEATURES

KEYWORDS SOURCE ORGANISM

REFERENCE AUTHORS TITLE JOURNAL

PEATURES

DEFINITION

RESULT 32 AX057492 LOCUS

ACCESSION VERSION

ô

Gaps

ô

PAT 17-JAN-2001

ö

Gaps

ö

PAT 17-JAN-2001

Gaps

ö

Indels

Length 20;

```
ch 0.5%; Score 20; DB 1; Similarity 100.0%; Pred. No. 40; 20; Conservative 0; Mismatches
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                    2658 AATTGGCAGGAAGCAACATC 2677
                                                                                                                                                     20 TTGGGACAACCAGCCTAAAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20 TTTATTGCACAGACTTGCGG 1
                                                                                                                                                                                                                                                                                                      AX057504.1 GI:12310238
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                       Homo sapiens (human)
                                                                                  Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Homo sapiens
                                                                                                                                                                                                                                                                                                                                                           Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
Best Local S
Matches 20
                                                                      Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 40
AX057509/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  DEFINITION
                                                                                    Best Loca
Matches
                                                                                                                                                                                                                                          AX057504/c
                                                                                                                                                                                                                                                                        DEFINITION
                                                                                                                                                                                                                                                                                                                                                           ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                          REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       JOURNAL
                                                                                                                                                                                                                                                                                         ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                             JOURNAL
                                                                                                                                                                                                                          RESULT 39
                                                                                                                                                                                                                                                                                                                         KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                            TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     LOCUS
                                                                                                                                                                                                                                                          LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 d
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                               PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalla, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                      Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Lorenz, E., Schwartz, D.A. and Schutte, B.C. variant tlr4 nucleic acid and uses thereof Patent: WO 0077204-A 35.21-DEC-2000; University of Iowa Research Foundation (US); Lorenz, Eva (US)
                                                                                         Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 33 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204.3 3 4 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ô
                                                                                                                                                                                                                                                                                             .;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Score 20; DB 1; Length 20;
Pred. No. 40;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         linear
                                                                                                                                                                                                                                                          Query Match 0.5%; Score 20; DB 1; Length 20; Best Local Similarity 100.0%; Pred. No. 40; Matches 20; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                 linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                 DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match 0.5%; Score 20; DB Best Local Similarity 100.0%; Pred. No. 40; Matches 20; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1. .20
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                               20 bp
Sequence 34 from Patent WO0077204.
AX057498
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20 bp
Sequence 35 from Patent WO0077204.
AX057499
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      2590 GACGACTCAGAAAAGCCCTG 2609
                                                                                                                                                                                                                                                                                                                              2467 TCATTGTCCTGCAGAAGGTG 2486
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      caccacrcacaaaaccccrc 20
                                                                                                                                                                                                                                                                                                                                                  1 TCATTGTCCTGCAGAAGGTG 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AX057499.1 GI:12310233
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AX057498.1 GI:12310232
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens (human)
Homo sapiens
        Homo sapiens (human)
Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Нощо варіепв
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                               DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           source
                                                                                                                                                                                 Bource
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
AUTHORS
          SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 38
                                                                                                                              JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 37
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       REFERENCE
                                                                                             AUTHORS
                                                                             REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AX057499
                                                                                                                                                                                                                                                                                                                                                                                                                                 AX057498
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          FEATURES
                                                                                                                                                                FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        임
                                                                                                                                                                                                                                                                                                                                  ð
```

```
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
    PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Bukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                      Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Lorenz,E., Schwartz,D.A. and Schutte,B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 45 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                    Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 40 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         .
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match 0.5%; Score 20; DB 1; Length 20; Best Local Similarity 100.0%; Pred. No. 40; Matches 20; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Score 20; DB 1; Length 20;
Pred. No. 40;
       linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DNA
         DNA

    .20
    /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   tch 0.5%; Score 20; DB al Similarity 100.0%; Pred. No. 40; 20; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                         /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20 bp
Sequence 45 from Patent WO0077204.
AXO57509
AXO57509.1 GI:12310243
20 bp
Sequence 40 from Patent WO0077204.
AX057504
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1351 TTGGGACAACCAGCCTAAAG 1370
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 733 TITATIGCACAGACTIGCGG 752
```

RESULT 41 AX057511/c LOCUS

DEFINITION ACCESSION VERSION KEYWORDS SOURCE

ORGANISM

REFERENCE AUTHORS

JOURNAL

source

FEATURES

```
PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Homo sapiens
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Butheria; Primates; Catarrhini; Hominidae; Homo.
                                                       Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 49 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tir4 nucleic acid and uses thereof
Patent: WO 0077204-A 50 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Lorenz,E., Schwartz,D.A. and Schutte,B.C.
Variant tir4 nucleic acid and uses thereof
Patent: WO 0077204-A 51 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 40; tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                Length 20;
                                                                                                                                                                                                                                                                                                      0; Indels
                                                                                                                                                                                                                                                              Query Match 0.5%; Score 20; DB 1; Best Local Similarity 100.0%; Pred. No. 40; Matches 20; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DNA
                                                                                                                                                            1. .20
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20 bp
Sequence 50 from Patent WO0077204.
AX057514.1 GI:12310248

    .20
    /organism="Homo sapiens"
/mol_type="unassigned DNA"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AX057515 20 bp
Sequence 51 from Patent WO0077204.
AX057515
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1947 AAGGACCAGAGGCAGCTCTT 1966
                                                                                                                                                                                                                                                                                                                                              1788 TGTCTGAACTCCCTCCAGGT 1807
                                                                                                                                                                                                                                                                                                                                                                        20 TGTCTGAACTCCCTCCAGGT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20 AAGGACCAGAGGCAGCTCTT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AX057515.1 GI:12310249
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
Best Local S:
Matches 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AX057515/c
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 44
AX057514/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ORGANISM
                                                                              TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ACCESSION
VERSION
                                      REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 45
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AUTHORS
                                                                                                                                         FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          LOCUS
                                                                                                                                                                                                                                                                                                                                                 à
                                                                                PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                        Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Buteleostomi,
Mammalia, Butheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                              (ns)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 48 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                               Lorenz,E., Schwartz,D.A. and Schutte,B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 47 21-DEC-2000;
University of Lowa Research Foundation (US); Lorenz, Eva
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   linear
                                                                                linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Length 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Score 20; DB 1;
Pred. No. 40;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DB 1;
                                                                                NA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
0.5%; Score 20; DB 3
Best Local Similarity 100.0%; Pred. No. 40;
Matches 20; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match 0.5%; Score 20; DB Best Local Similarity 100.0%; Pred. No. 40; Matches 20; Conservative 0; Mismatches

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

    20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                        20 bp
Sequence 47 from Patent WO0077204.
AX057511
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AX057513 20 bp
Sequence 49 from Patent WO0077204.
AX057513. GI:12310247
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 48 from Patent WO0077204.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1656 TTGACCTTCCTGGACCTCTC 1675
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1568 IGGCTIGICCAGICTCGAAG 1587
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              20 TGGCTTGTCCAGTCTCGAAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AX057512.1 GI:12310246
                                                                                                                                       AX057511.1 GI:12310245
                                                                                                                                                                             Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Homo sapiens (human)
Homo sapiens
                                                                                                                                                                                                     Homo sapiens
```

RESULT 42 AX057512/c LOCUS

셤

DEFINITION

ACCESSION VERSION

ORGANISM

KEYWORDS SOURCE

REFERENCE AUTHORS

JOURNAL

TITLE

FEATURES

ö

ACCESSION VERSION KEYWORDS SOURCE ORGANISM

AX057513/c LOCUS DEFINITION

RESULT 43

PAT 17-JAN-2001

10001863-3.81.rge

ठ 쉽

```
Homo sapiens (human)
Homo sapiens
Eukaryota, Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                iorenz,E., Schwartz,D.A. and Schutte,B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 007204-A 54 Z1-DEC-2000;
University of Iowa Research Foundation (US) ; Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Lorenz,E., Schwartz,D.A. and Schutte,B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: Wo 0077204-A 55 21-DEC-2000;
University of Iowa Research Foundation (US) ; Lorenz, Eva
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0; Indels
                                                                                                                                                                                                                                                                                                                                                                               0.5%; Score 20; DB 1; Length 20;
00.0%: Pred. No. 40;
                             linear
                                                                                                                                                                                                                                                                                                                                                                                                                0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.5%; Score 20; DB 1;
100.0%; Pred. No. 40;
tive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DNA
                               DNA
                                                                                                                                                                                                                                                                                                                                                                                 Query Match 0.5%; Score 20; UB Best Local Similarity 100.0%; Pred. No. 40; Matches 20; Conservative 0; Mismatches

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

    .20
    /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20 bp
Sequence 56 from Patent WO0077204.
AX057520
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20 bp
Sequence 55 from Patent WO0077204.
AX057519
                     20 bp
Sequence 54 from Patent WO0077204.
AXO57518
AXO57518.1 GI:12310252
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  2484 GIGGAGAAGACCCIGCICAG 2503
                                                                                                                                                                                                                                                                                                                                                                                                                                                     2380 AGCACTTCATCCAGAGCCGC 2399
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20 GTGGAGAGACCCTGCTCAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20 AGCACTTCATCCAGAGCCGC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AX057520.1 GI:12310254
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AX057519.1 GI:12310253
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens (human)
                                                                                                                 Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens
                                                                                                                                     Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DEFINITION
ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                source
                                             DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                 source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT 50
AX057520/c
            AX057518/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 49
AX057519/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  SOURCE
ORGANISM
                                                                                                                     SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Matches
                                                                                                                                                                                                                         TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ACCESSION
VERSION
                                                                                                                                                                                       REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            FEATURES
                                                                                                                                                                                                                                                                             FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Locus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                           Š
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          PAT 17-JAN-2001
                                                                     ö
                                                                                                                                                                                                                           PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                                                                                                  Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalla, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 53 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                  Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 52 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DB 1; Length 20; 40;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match 0.5%; Score 20; DB 1; Length 20; Best Local Similarity 100.0%; Pred. No. 40; Matches 20; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0; Indels
                                  Length 20;
                                                                                                                                                                                                                             linear
                                                                   0; Indels
                                  DB 1;
                                                                                                                                                                                                                             DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1. .20
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match 0.5%; Score 20; DB Best Local Similarity 100.0%; Pred. No. 40; Matches 20; Conservative 0; Mismatches
                                ch 0.5%; Score 20; DB 1 Similarity 100.0%; Pred. No. 40; 20; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20 bp
Sequence 53 from Patent WO0077204.
AX057517
                                                                                                                                                                                                                             20 bp
Seguence 52 from Patent WO0077204.
AX057516
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               2142 CTTCTTGCTGCTGCATAAA 2161
                                                                                                      2074 CGGTCCTCAGTGTGCTTGTA 2093
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           CTTCTTGCTGGCTGCATAAA 1
                                                                                                                          20 CGGTCCTCAGTGTGCTTGTA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AX057517.1 GI:12310251
                                                                                                                                                                                                                                                                                  AX057516.1 GI:12310250
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                   Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                       Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20
                                      Query Match
Best Local &
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AX057517/c
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                   KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                source
                                                                                                                                                                                                                             LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                            RESULT 46
AX057516/c
                                                                                                                                                                                                                                                                                                                                                                                         REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        REFERENCE
AUTHORS
                                                                     Matches
                                                                                                                                                                                                                                                                ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                            JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 47
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    SOURCE
```

셤 ð

ö

Gaps

; 0

PAT 17-JAN-2001

PAT 17-JAN-2001

à

ö

Gaps

ö

 $(\Omega S)$ 

Gaps

ö

0; Indels

PAT 17-JAN-2001

linear

DNA

```
0.5%; Score 20; DB 1;
                 100.0%; Pred. No. 40; tive 0; Mismatches
                                                                                                                                                                      AX057524 20 bp
Sequence 60 from Patent WO0077204.
AX057524 GI:12310258
                                                                2725 CTGGGTCCAACACTTGTTCA 2744
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 448 ATGGGGCATATCAGAGCCTA 467
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1 GCTCACAGAAGCAGTGAGGA 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                15 GCTCACAGAAGCAGTGAGGA 34
                                                                                 CTGGGTCCAACACTTGTTCA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20 ATGGGGCATATCAGAGCCTA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AX057525.1 GI:12310259
                                                                                                                                                                                                                                                  Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens (human)
                 1 Similarity 100.
20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Similarity
                                                                                                                                                                                                                                                                    Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
Best Local Simi]
Matches 20; (
   Query Match
Best Local S
Matches 20
                                                                                             20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            source
                                                                                                                                                                                                                                                SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 54
AX057525/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   SOURCE
ORGANISM
                                                                                                                                                                                     DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ACCESSION
VERSION
                                                                                                                                                                                                     ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                             REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 55
AX752002
                                                                                                                                                                                                                                                                                                                                                          JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             JOURNAL
                                                                                                                                          RESULT 53
AX057524
                                                                                                                                                                                                                                   KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                          FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               TITLE
                                                                                           g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            셤
                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                             PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               PAT 17-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens
Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                    Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A SS 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US).
Location/Qualifiers
              Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 56 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 57 21-DEC-2000;
University of Iowa Research Foundation (US) ; Lorenz, Eva (US)
                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                      Length 20;
                                                                                                                                                                                                                                                                                                                                             linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Length 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      0; Indels
                                                                                                                                                                     0.5%; Score 20; DB 1;
100.0%; Pred. No. 40;
ative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.5%; Score 20; DB 1;
                                                                                                                                                                                                                                                                                                                                             DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DNA
                                                                                           1. .20
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     100.0%; Pred. No. 40; ive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1. .20
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                       20 bp
Sequence 57 from Patent WO0077204.
AX057521
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AX057522 20 bp
Sequence 58 from Patent WO0077204.
                                                                                                                                                                                                                                2659 ATTGGCAGGAAGCAACATCT 2678
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2590 GACGACTCAGAAAAGCCCTG 2609
                                                                                                                                                                                                                                                     20 ATTGGCAGGAAGCAACATCT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       20 GACGACTCAGAAAAGCCCTG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AX057522
AX057522.1 GI:12310256
                                                                                                                                                                                                                                                                                                                                                                                         AX057521.1 GI:12310255
                                                                                                                                                                                                                                                                                                                                                                                                                       Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Homo sapiens (human)
                                                                                                                                                                  Query Match
Best Local Similarity 100.0
Matches 20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                       Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Best Local Similarity
Matches 20; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
                                                                                                                                                                                                                                                                                                                                                     DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 52
AX057522/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        DEFINITION
ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       KEYWORDS
SOURCE
ORGANISM
                                                                                             source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              source
                                                                                                                                                                                                                                                                                                                                                                                                                      SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                          AX057521/c
REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  REFERENCE
AUTHORS
                                            JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               JOURNAL
                                                                                                                                                                                                                                                                                                             RESULT 51
                                                                           FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            PEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              FEATURES
                                                                                                                                                                                                                                                                                                                                            LOCUS
                                                                                                                                                                                                                                  ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ð
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             PAT 17-JAN-2001
Eukaryota; Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                               Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 60 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tir4 nucleic acid and uses thereof
Patent: WO 0077204-A 61 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Butel
Mammalia, Butheria, Primates, Catarrhini, Hominidae, Homo
                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                               Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Score 20; DB 1; Length 20;
Pred. No. 40;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             linear
                                                                                                                                                                                                                                                                                                                                           Indele
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0; Indels
                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                               ï
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0.5%; Scor.
100.0%; Pred. No. 30,
... 0; Mismatches

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                               0.5%; Score 20; DB ilarity 100.0%; Pred. No. 40; Conservative 0; Mismatches

    .20
    /organism="Homo sapiens"
    /mol type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 61 from Patent WO0077204. AX057525
```

PAT 17-JAN-2001

```
Homo sapiens
Homo sapiens
Bukaryota; Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi;
                                                                                                                                                 Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                           Lorenz, E., Schwartz, D.A. and Schutte, B.C. variant tlr4 nucleic acid and uses thereof patent: WO 0077204-A 18 21-DEC-2000; University of Iowa Research Foundation (US); Lorenz, Eva (US) Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Arditi,M., Rajavashisth,T. and Shah,P.K.
Treating vascular disease by inhibiting Toll-like receptor-4
Patent: WO 03035110-A 4 01-MAY-2003;
CEDARS-SINAI MEDICAL CENTER (US)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0.5%; Score 19; DB 1; Length 19;
00.0%; Pred. No. 45;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Indels
                                                                                                                                                                                                                                                                                                                                                      Length 19;
                                         linear
                                                                                                                                                                                                                                                                                                                                                                                         ö

    .19
        /organism="synthetic construct"
/mol_type="unassigned DNA"
/db_xref="taxon:32630"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        DNA
                                                                                                                                                                                                                                                                                                                                                        0.5%; Score 19; DB 1;
100.0%; Pred. No. 45;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      other sequences; artificial sequences.
                                          DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Ouery Match 0.5%; Score 19; DB
Best Local Similarity 100.0%; Pred. No. 45;
Matches 19; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 18 bp
Sequence 26 from Patent WO0077204.
AXO57490
AXO57490.1 GI:12310224
                                                                                                                                                                                                                                                                                                                                                                       larity 100.0%; Pred. No. 45; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                              1. .19
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AX752003 19 bp
Sequence 4 from Patent W003035110.
AX752003 AX752003.1 GI:32134126
                                     19 bp
Sequence 18 from Patent WO0077204.
AX057482
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              2256 GAAGGGGTGCCTCCATTTC 2274
                                                                                                                                                                                                                                                                                                                                                                                                                        812 CITIAICCAACCAGGIGCA 830
                                                                                                                                                                                                                                                                                                                                                                                                                                            CTTTATCCAACCAGGTGCA 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /note="Primer"
                                                                                          AX057482.1 GI:12310216
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               GAAGGGTGCCTCCATTTC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              synthetic construct
synthetic construct
                                                                                                                      Homo sapiens (human)
Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                            l Similarity
19; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             19
                                                                                                                                                                                                                                                                                                                                                             Query Match
                                                                                                                                                                                                                                                                                                                                                                               Local
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DEFINITION
ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  LOCUS
                                                                                                                                                                                                                                                                                   source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      RESULT 58
AX752003/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RESULT 59
AX057490
LOCUS
                                                           DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AUTHORS
TITLE
JOURNAL
                                                                                                                                          ORGANISM
                                                                                                                                                                                     REFERENCE
AUTHORS
TITLE
              RESULT 57
AX057482
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                           Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             REFERENCE
                                                                                                                                                                                                                                   JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         FEATURES
                                                                                                                                                                                                                                                                  FEATURES
                                                                                                                             SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               OS Artificial Sequence
PN W0 017293-A/6
PD 04-0CT-2001
PF 02-APR-2001 WO 2001JP002869
PR 31-MAR-2000 JP 00P 99617,22-NOV-2000 JP 00P 356719 PR
28-WAR-2001 US 99615-8
PI SHOJI FURUSAKO,SADAO MORI,KAMON SHIRAKAWA,TOMOHIRO TAKAHASHI
PC C12M15/00,C07K7/08,C07K14/705,C07K16/28,A61K45/00,A61P31/04,
PC A61P39/395,A61P43/00,G01N33/15,G01N33/50,G01N33/577
PC TLR/CD14 binding inhibitor
PC TLR/CD14 binding inhibitor
FH Key 1. .20
FT SOURCE 1. .20
//organism='Artificial Sequence'.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                 ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            PAT 27-AUG-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 PAT 20-JUN-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                          Arditi,M., Rajavashisth,T. and Shah,P.K.
Treating vascular disease by inhibiting Toll-like receptor-4
Patent: WO 03035110-A 3 01-MAY-2003;
CEDARS-STAIR MEDICAL CENTER (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                   .;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /organism='Artificial Sequence'. Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ch 0.5%; Score 20; DB 1; Length 20; Similarity 100.0%; Pred. No. 40; 20; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                  Score 20; DB 1; Length 20;
Pred. No. 40;
                                                                                                                                                                                                                                                                                                                                                   Indels
      linear

    .20
    /organism="synthetic construct"
/mol_type="genomic DNA"
    /db_xref="taxon:32630"

                                                                                                                                                                                                                            1. .20
/organism="synthetic construct"
/mol_type="unassigned DNA"
/db_xref="taxon:32630"
/noTe="Primer"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DNA
      DNA
                                                                                                                other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                    0.5%; Score 20; DB
llarity 100.0%; Pred. No. 40;
Conservative 0; Mismatches
20 bp
Sequence 3 from Patent WO03035110.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20 bp
                                                                                                                                                                                                                                                                                                                                                                                  1768 TGGATACGTTTCCTTATAAG 1787
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             501 CCCATCCAGAGTTTAGCCCT 520
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               BD102553
TLR/CD14 binding inhibitor.
BD102553
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1 recaracerricerrarase 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          BD102553.1 GI:22648127
WO 0172993-A/6.
                                                    AX752002.1 GI:32134125
                                                                                    synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                    Local Similarity
nes 20; Conserva
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
Best Local Similarity
Matches 20; Conserv
                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           source
                                                                     KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ORGANISM
                                                                                                                                                                                                                                                                                                                                         Best Loca
Matches
                        DEFINITION
                                                                                                                                                               TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 56
BD102553
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     TITLE
                                                                                                                                  REFERENCE
AUTHORS
                                       ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          FEATURES
                                                                                                                                                                                                               FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     COMMENT
                                                                                                                                                                                                                                                                                                                                                                                                                  요
                                                                                                                                                                                                                                                                                                                                                                                       ò
```

PAT 20-JUN-2003

ö

Gaps

.. 0

PAT 17-JAN-2001

ö

Gaps

Gaps

ö

PAT 25-SEP-2002

BD161176/c

RESULT 60

δ 셤 ORGANISM

REFERENCE AUTHORS

TITLE

JOURNAL

COMMENT

FEATURES

ACCESSION VERSION KEYWORDS SOURCE

REFERENCE AUTHORS

JOURNAL

FEATURES

```
Dergeron, M.G., Picard, F.J., Ouellette, M. and Roy, P.H.
Bergeron, M.G., Picard, F.J., Ouellette, M. and Roy, P.H.
Species-specific and universal DNA probes and amplification primers to rapidly detect and identify common bacterial pathogens and associated antibiotic resistance genes from clinical specimens for routine diagnosis in microbiology laboratories
Patent: US 5994666-A 219 30-NOV-1999;
Location/Qualifiers
                                                                          Antisense oligonucleotide modulation of B-raf gene expression
Patent: US 5981731-A 25 09-NOV-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Antisense oligonucleotide inhibition of raf gene expression Patent: US 6410518-A 114 25-JUN-2002; Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 DB 1; Length 20;
                                                                                                                                                                                                          Query Match 0.4%; Score 17; DB 1; Length 20; Best Local Similarity 100.0%; Pred. No. 76; Matches 17; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                             linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Query Match 0.4%; Score 17; DB 1;
Best Local Similarity 100.0%; Pred. No. 76;
Matches 17; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                               DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DNA
                                                                                                                                                                                                                                                                                                                                                                                                                     20 bp DN Sequence 114 from patent US 6410518. AR216067
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20 bp DN Sequence 219 from patent US 5994066. AR089460
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.4%; Score 16.8; D
90.0%; Pred. No. 79;
tive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                      /mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /mol_type="genomic DNA"
                                                                                                                Location/Qualifiers
                                                                                                                                                  organism="unknown"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             organism="unknown"
                                                                                                                                                                                                                                                                                        1465 TGAAACAAATGAGTGAG 1481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1465 TGAAACAAATGAGTGAG 1481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AR216067.1 GI:23314355
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AR089460.1 GI:10016217
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20 TGAAACAAATGAGTGAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1 (bases 1 to 20).
Monia, B.P.
                  Unclassified.
1 (bases 1 to 20)
Monia, B.P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
Best Local Similarity 90.0
Matches 18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 source
                                                                                                                                      Bource
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         source
                                                                                                                                                                                                                                                                                                                                                                                     RESULT 62
AR216067/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 63
AR089460/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    SOURCE
ORGANISM
  ORGANISM
                                                                            TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                             DEFINITION
                                      REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               REFERENCE
                                                                                                                FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                VERSION
                                                                                                                                                                                                                                                                                          à
                                                                                                                                                                                                                                                                                                                            셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                   22 bp DNA linear PAT 17-JAN-2003 Protein participating in mechanism for preventing infection and gene encoding the protein.
                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         PAT 01-SEP-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        C (C12NS/10,C12R1:91),C12N1S/00,C12N5/00,(C12N5/00,C12R1:91) CC Description of Artificial Sequence: PCR primer FH Key Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               other sequences; artificial sequences.

(Dases 1 to 22)

Yoshikat,Y., Matsuguchi,T. and Iwami,K.
Protein participating in mechanism for preventing infection and gene encoding the protein
Patent: JP 2002176986-A 14 25-UUN-2002;
NAGOYA INDUSTRIAL SCIENCE RESEARCH INSTITUTE
OS Artificial Sequence
PN JP 2002176986-A/14
                                  Lorenz, E., Schwartz, D.A. and Schutte, B.C.
Variant tlr4 nucleic acid and uses thereof
Patent: WO 0077204-A 26 21-DEC-2000;
University of Iowa Research Foundation (US); Lorenz, Eva (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            PD 25-JUN-2002
PF 14-DEC-2000 JP 2000380561
PI YASUNOBU YOSHIKAL, TETSUYA MATSUGUCH1, KENICHIRO IWAMI PC C12N15/09, C07K14/705, C12N1/15, C12N1/19, C12N1/21, C12N5/10// PC
Mammalia; Butheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                                                                                    ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ce 1. .22
/organism='Artificial Sequence'
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        DB 1; Length 22;
                                                                                                                                                                                                                              DB 1; Length 18;
51;
                                                                                                                                                                                                                                                                    0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       linear

    .22
        /organism="synthetic construct"
/mol_type="genomic DNA"
        /db_xref="taxon:32630"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DNA
                                                                                                                                                                                                                            Query Match

0.5%; Score 18; DB
Best Local Similarity 100.0%; Pred. No. 51;
Matches 18; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.5%; Score 17.2; D
86.4%; Pred. No. 84;
tive 0; Mismatches
                                                                                                                                                    /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 25 from patent US 5981731.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2225 GGTAAGGAATGAGCTAGTAAAG 2246
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         GCTGAGAATGAGCTGGTAAAG 1
                                                                                                                                                                                                                                                                                                        TTCATTGGATACGTTTCC 1780
                                                                                                                                                                                                                                                                                                                                              rrcarregaracerrrcc 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       BD161176.1 GI:27866934
JP 2002176986-A/14.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 GI:10012314
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             19; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Best Local Similarity
Matches 19; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AR085547.1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AR085547
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AR085547
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          C12P21/
                                                                                                                                                                                                                                                                                                          1763
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         22
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                      source
                                                                                                                                                                                                                                                                                                                                                                                                                                         LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              source
```

ö

Gaps

ö

PAT 07-SEP-2000

ö

Gapa

ö

RESULT 61 AR085547/c

ઠે 셤 Locus

ò 셤

```
PAT 16-JUN-2001
                                                                                                                                                                                                          PAT 20-DEC-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            PAT 14-FEB-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1 (bases 1 to 21)
Mezes, P.S., Richard, R.A., Affholter, J.A. and Kotite, N.J.
Dimer and multimer forms of single chain polypeptides
Patent: US 6071515-A 87 06-JUN-2000;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                               ö
                               Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Length
                                                                                                                                                                                                                                                                                                                        Unclassified.

1 (bases 1 to 20)
Ward, D.T. and Watt, A.T.
Antisense modulation of helicase-moi expression
Patent: US 6444466-A 14 03-SEP-2002;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    3,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Match 0.4%; Score 16.2; DB 1; Local Similarity 85.7%; Pred. No. 96; Les 18; Conservative 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DB 1;
                                 DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       21 bp DNA
Sequence 108 from patent US 6197505.
AR137361.1 GI:14478870
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DNA
                               Score 16.4; DE Pred. No. 86; 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      21 bp 1
Sequence 87 from patent US 6071515.
AR097024
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1. .21
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0.4%; Score 16.4; larity 94.4%; Pred. No. 86; Conservative 0; Mismatches
                                                                                                                                                                                              AR226133 20 bp
Sequence 14 from patent US 6444466.
AR226133
                                                                                                                                                                                                                                                                                                                                                                                                                                               /mol_type="genomic DNA"
/db_xref="taxon:32644"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    2683 GAAGAGAAAAATAAAAACCT 2703
                                                                                                                                                                                                                                                                                                                                                                                                                         1. .20
/organism="unknown"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     21 GATGATGACAAATAAAAACCT 1
                                                                                               1640 CACAGAGCTGAGAAACTT 1657
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   613 AAACTTTGAAAGAACTTA 630
                                                                                                                               CACAGAGCTGAGAACAT 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AR097024.1 GI:12805754
                                                                                                                                                                                                                                                          AR226133.1 GI:27264287
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    19 AAACTCTGAAAGAACTTA 2
                               0.4%;
ilarity 94.4%;
Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Unknown.
Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Best_Local Similarity
Matches 17; Conserv
                                                  Local Similarity
                                                                                                                                                                                                                                                                                          Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Unknown.
                                                                                                                                                                                                                                                                                                           Unknown.
                                                                  17;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
Best Local Si
Matches 18;
                                   Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DEFINITION
ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             source
                                                                                                                                                                                                                                                                                                                                                                                                                           source
                                                                                                                                                                               RESULT 66
AR226133/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AR137361/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                DEFINITION
                                                                                                                                                                                                                              DEFINITION
                                                                                                                                                                                                                                                                                                            ORGANISM
                                                                                                                                                                                                                                                                                                                                                        AUTHORS
TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AR097024/C
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ACCESSION
VERSION
KEYWORDS
                                                     Best Loc
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 67
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      RESULT 68
                                                                                                                                                                                                                                             ACCESSION
                                                                                                                                                                                                                                                                                                                                         REFERENCE
                                                                                                                                                                                                                                                                           KEYWORDS
SOURCE
                                                                                                                                                                                                                                                              VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ð
                                                                                                                               셤
                                                                                                                                                                                                                                                                                                      Streptococcus.

Streptococcus.

I (bases 1 to 20)

Series - Patentd, F.G., Weretto, M. and Roy, P.H.

Species - Specific, genus-specific and universal probes and primers for quickly detecting and identifying common bacterial and fungal pathogens and relating antibiotic tolerance genes from clinical specimens for diagnosis in microbiological laboratory

I NPECTIO DIAGNOSTICS INC

PR 07-ARR-2001

PR 03-ARR-2001

PR 04-NOV-1995 US 08/743637

PI MICHEL UU BERGERON, FRANCOIS G PICARD, MARC WERETTO, PAUL H ROY

PO CILILS/09, CILINI/21, CIL2Q1/68, CILZR1:01), (CIL2Q1/68, PC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          PAT 21-JAN-2000
                                                                                                                                  PAT 27-AUG-2002
                                                                                                                               BD022993
Species-specific, genus-specific and universal probes and primers for quickly detecting and identifying common bacterial and fungal pathogens and relating antibiotic tolerance genes from clinical specimens for diagnosis in microbiological laboratory.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (11201/68, C12R1:44), (C12Q1/68, C12R1:72), C12N15/00 CC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                               Sacteria; Firmicutes; Lactobacillales; Streptococcaceae;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Matthijs,G.
CARBOHYDRATE-DEFICIENT GLYCOPROTEIN SYNDROME TYPE I
Patent: WO 9849324-A 24 05-NOV-1998;
MATHLIG GERT (BE); GENZYME LTD (GB)
LOCATION/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Indels

    .20
    forganism="Streptococcus salivarius"
/mol type="genomic DNA"
    /db_xref="taxon:1304"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 2;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1 Similarity 90.0%; Pred. No. 79; 18; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1. .20
/organism="unidentified"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%; Score 16.8; 1
00.0%; Pred. No. 79;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20 bp
Sequence 24 from Patent WO9849324,
A83595
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                2378 CCAGCACTTCATCCAGAGCC 2397
                       CCAGCACTICATCCAGAGCC 2397
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      CCAGCACTTCATCAAGAGTC 1
                                                                                                                                                                                                                                               JP 2001504330-A/61.
Streptococcus salivarius
Streptococcus salivarius
                                                     20 ccagcacricarcaacacrc 1
                                                                                                                                                                                                                                   BD022993.1 GI:22564216
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Strandedness: Single;
CC Topology: Linear;
FH Key L
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          A83595.1 GI:6732851
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (bases 1 to 20)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          C12R1:46)
                                                                                                                                                                                                                    BD022993
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ы
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20
                       2378
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                      BD022993/c
                                                                                                                                     LOCUS
                                                                                                                                                                                                                                                                                ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               RESULT 65
A83595
                                                                                                                                                                                                                                                                                                                                               AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          REFERENCE
                                                                                                                                                                                                                  ACCESSION
                                                                                                                                                                                                                                                                                                                                REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                              JOURNAL
                                                                                                                                                                                                                               VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                              COMMENT
                                                                                                     RESULT
```

셤 ð

ö

PAT 24-MAR-2004

linear

```
Sendtner,M. and Boemmel,H.
Test system for the discovery of active agents in nerve cell
                                                                                                                                                                                                   Patent: WO 2004020661-A 11 11-MAR-2004;
Medlnnova Gesellschaft fuer Medizinische Innovation en aus
Akademischer Forschung mbH (DE)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ce 1. .21
/organism='Artificial Sequence'.
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                           DB 1; Length 21;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DB 1; Length 21;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                          3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            21 bp DNA 11:
Method for detecting Kawasaki disease factor.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Method for detecting Kawasaki disease factor
Patent: JP 2000157297-A 20 13-JUN-2000;
SHIONOGI & CO LTD
OS Artificial Sequence
PN JP 2000157297-A/20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               TAKESHI YOSHIOKA,RYUJI SUZUKI
C12Q1/68,C12N15/09,G01N33/48,C12N15/00
                                                                                                                                                                                                                                                                                        /organism="synthetic construct"
/mol type="unassigned DNA"
/db_xref="taxon:32630"
/noce="primer"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /organism="synthetic construct"
/mol_type="genomic DNA"
/db_xref="taxon:32630"
21 bp DN/
Sequence 11 from Patent WO2004020661.
CQ787005
                                                                                                      synthetic construct
other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           other sequences; artificial sequences.
1 (bases 1 to 21)
Yoshioka,T. and Suzuki,R.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ; Score 16.2; DE; Pred. No. 96; 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                          0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                         0.4%; Score 16.2;
85.7%; Pred. No. 96;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                01-DEC-1998 JP 1998341661
                                                                                                                                                                                                                                                                                                                                                                                                                                           342 GACCTGAGCTTTAATCCCCTG 362
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 409 TGCTGGATTTATCCAGGTGTG 429
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          21 GACCTGAGTTTGAATCCCCAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      CQ787005.1 GI:45721988
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           E35929.1 GI:18624640 JP 2000157297-A/20.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match 0.4%;
Best Local Similarity 85.7%;
Matches 18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            synthetic construct
synthetic construct
                                                                                      synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                          18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                13-JUN-2000
                                                                                                                                                                                                                                                                                                                                                                                         Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  source
                                                                                                                                                                                       diseases
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             E35929
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               OS DE PROPERTIES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  21
                                                                                                                                                                                                                                                                                                                                                                           Query Match
     LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                            source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                       Matches
                                                                                                                                                    AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 71
E35929/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           TITLE
JOURNAL
                                                                                                                                     REFERENCE
                                                                                                                                                                                                     JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 72
                                                                                                                                                                                                                                                       FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               COMMENT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       .
G
                                                                                                                                                                                                                                                                                                                                                                                                                                             ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              셤
                                                                                                                                                                                                                                                                                                                                                                                                                                          BD231344 217-JUL-2003 Genes for assessing cardiovascular status and compositions for use
                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        other sequences; artificial sequences.
I toase I to 2.1
Norberg.L.T., Anderson.M.K., Lindstrom,P.H.R. and Jonsson,L.
Genes for assessing cardiovascular status and compositions for use
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Genes
                                                                 1 (bases 1 to 21)
Norberg,L.Torbjorn., Andersson,M.Kristina. and
Lindstrom,P.Harry.Rutger.
Methods for assessing cardiovascular status and compositions for use thereof
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PI
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      PI LENA JONSSON
PC C12Q1/68,124-OCT-1998 US 60/104302
PI LENA JONSSON
PC C12Q1/68,12N15/09//G01N33/53,G01N33/566,C12N15/00 CC GE for assessing cardiovascular status and compositions for Use thereof
PC Use thereof
PH Key LOCAF***
FT SOURCE
                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         PAIROSEAKENSINGU AB
OS Artificial Sequence
PN JP 2002527079-A/108
PD 27-AUG-2002
PF 13-AUG-1999 JP 2000576056
PR 14-OCT-1999 US 60/104286,14-OCT-1998 US 60/104302
LEIF TORBJORN NORBERG,MARIA KRISTINA ANDERSSON, PER HARRY PI
                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                       DB 1; Length 21;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 DB 1; Length 21;
                                                                                                                                                                                                                                                                                        IndelB
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Patent: JP 2002527079-A 108 27-AUG-2002;

    21
    ^2ynthetic construct"
/mol_type="genomic DNA"
/db_xref="taxon:32630"

                                                                                                                                                 Patent: US 6197505-A 108 06-MAR-2001;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match 0.4%; Score 16.2; Di
Best Local Similarity 85.7%; Pred. No. 96;
Matches 18; Conservative 0; Mismatches
                                                                                                                                                                                                                                                     0.4%; Score 16.2; D
85.7%; Pred. No. 96;
iive 0; Mismatches

    21
    /organism="unknown"
    /mol_type="unassigned DNA"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Location/Qualifiers
                                                                                                                                                                                                                                                                                                                       27 AGTGAGGATGATGCCAGGATG 47
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 27 AGTGAGGATGATGCCAGGATG 47
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 21 AGTGAGGTTGATGCCAGAAG 1
                                                                                                                                                                                                                                                                                                                                                      21 AGTGAGGTTGATGCCAGAAAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        BD231344.1 GI:33041114 JP 2002527079-A/108.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                          Conservative
                                                   Unclassified.
                                                                                                                                                                                                                                                  Query Match
Best Local Similarity
Matches 18; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          thereof.
                 Unknown.
                                    Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                       source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   source
                                                                                                                                                                                                                                                                                                                                                                                                                       BD231344/c
                                                                                                                                                                                                                                                                                                                                                                                                                                          LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 70
CQ787005/c
                                  ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        REFERENCE
AUTHORS
TITLE
                                                                 REFERENCE
AUTHORS
                                                                                                                                                   JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 69
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           JOURNAL
   KEYWORDS
                                                                                                                                                                    PEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                FEATURES
                                                                                                                    TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            COMMENT
                                                                                                                                                                                                                                                                                                                                                     셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ò
```

ö

Gaps

ö

PAT 31-JAN-2002

linear

ö

Gaps

Gaps

```
ABU86505 21 bp DNA linear SYN 21-MAY-2003 Synthetic construct DNA, forward primer for Japanese flounder microsatellite sequence PolillITUF.
                                                                                                                                                                                                                                                                                                                                                                                PAT 27-AUG-2002
                                                                                                                                                                                                                                                                                                                                                                            21 bp DNA linear PAT 27-AUG-20 Methods for assessing cardiovascular status and compositions for use thereof. BD075235
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           synthetic construct
synthetic construct
other sequences; artificial sequences.
I (bases 1 to 2).
Norberg.L.T., Andersson, M.K. and Lindstrom, P.H.R.
Methods for assessing cardiovascular status and compositions for use thereof
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Patent: JP 2001519660-A 108 23-OCT-2001;

Patent: JP 2001519660-A 108 23-OCT-2001;

EURONA MEDICAL AB

OS Artificial Sequence
PN JP 2001519660-A/108
PP 01-APR-1998 US 60/042930
PP 01-APR-1997 US 60/042930
PP ILET TORBORN NORBERG, MARIA KRISTINA ANDERSSON, PER HARRY
RUTGER LINDSTROM
PC C1201/68, COTX14/72, COTX14/575, C12N9/48
PC C1201/68, COTXI4/72, COTXI4/575, C12N9/48
PC Description of Artificial Sequence: PCR PRIMER FH KANY
; GEMINI GENOMICS AB (SE) ; LINDSTROM PER HARRY RUTGER (SE) SANDERS RHIANNON (SE) LOCATION/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match 0.4%; Score 16.2; DB 1; Length 21; Best Local Similarity 85.7%; Pred. No. 96; Aatches 18; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           C1201/68,C07K14/72,C07K14/575,C12N9/48
Description of Artificial Sequence: PCR PRIMER
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              /organism='Artificial Sequence'
                                                                                                                                                                         Length 21;
                                                                                                                                                                                                             Indels
                                                                                                                                                                     ch 0.4%; Score 16.2; DB 1; Similarity 85.7%; Pred. No. 96; 18; Conservative 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /organism="synthetic construct"
/mol type="genomic DNA"
/db_xref="taxon:32630"
                                                                                /organism="synthetic construct"
/mol type="unassigned DNA"
/mol type="taxon:32630"
/note="Oligonucleotide primer"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               synthetic construct
synthetic construct
other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 27 AGTGAGGATGATGCCAGGATG 47
                                                                                                                                                                                                                                                    27 AGTGAGGATGATGCCAGGATG 47
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        21 AGTGAGGTTGATGCCAGAAAG 1
                                                                                                                                                                                                                                                                                    21 AGTGAGGTTGATGCCAGAAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AB086505
AB086505.1 GI:28804357
                                                                                                                                                                                                                                                                                                                                                                                                                                                         BD075235.1 GI:22620838
JP 2001519660-A/108.
                                                                                                                                                                                                 Best Local Similarity
Matches 18; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              source
                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AB086505/c
                                                                                                                                                                                                                                                                                                                                                   RESULT 75
BD075235/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                      DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     RESULT 76
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                             ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                             VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   FEATURES
                                                   FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          COMMENT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sanders, R. Genetic polymorphic pattern for assessing disease Genetic polymorphism and polymorphic patterns, and compositions for use thereof Patent: WO 0055922-A 108 28-SEP-2000; NORBERG LEIF TORBJORN (SE); OLAISSON ERIK (SE); JONSSON LENA (SE)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PAT 16-NOV-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 PAT 12-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                 PAT 20-DEC-2002
                                                                                                                                                                                                           Gatti,R.A.
Modified SSCP method using sequential electrophoresis of multiple
nucleic acid segments
Patent: US 6458536-A 75_01-OCT-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Norberg, L.T., Olaisson, E., Jonsson, L., Lindstrom, P.H. and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Unclassified.

1 (bases 1 to 21)
Mezea, P.S., Richard, R.A., Affholter, J.A. and Kotite, N.J.
Dimer and multimer forms of single chain polypeptides
Patent: US 6329507-A 87 11-DEC-2001;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
0.4%; Score 16.2; DB 1; Length 21;
Best Local Similarity 85.7%; Pred. No. 96;
Matches 18; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                        Length 21;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              linear
                                                                                                                                                                                                                                                                                                                                                                                                              3; Indels
                                     linear
                                                                                                                                                                                                                                                                                                                                                                            DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          21 bp
Sequence 108 from Patent WO0056922.
AX037483
                                                                                                                                                                                                                                                                                                                                                                          Query Match 0.4%; Score 16.2; D
Best Local Similarity 85.7%; Pred. No. 96;
Matches 18; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AR367361 21 bp 1
Sequence 87 from patent US 6329507.
AR367361, GI:34600438
                                       21 bp | |
Sequence 75 from patent US 6458536.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1. .21
/organism="unknown"
/mol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                     1. .21
/organism="unknown"
/mol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                2683 GAAGAGAAAATAAAAACCT 2703
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Location/Qualifiers
                                                                                                                                                                                                                                                                                  Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                   1842 AAAAAACAGGAACTACAGCAT 1862
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1 AAAAACAGGAAGAACAGGAT 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        21 GATGATGACAAATAAAAACCT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AX037483.1 GI:11226910
                                                                                            AR233713.1 GI:27276337
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             synthetic construct
synthetic construct
                                                                                                                                                                                         (bases 1 to 21)
                                                                                                                                                       Unknown.
Unclassified.
                                                                                                                                  Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AX037483/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                     AR233713
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                         source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AR367361/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ORGANISM
                                                                                                                                  SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 JOURNAL
                                                                          ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 74
                                                                                                                                                                                                                                                                    JOURNAL
                                                                                                                                                                                                               AUTHORS
                                                                                                                                                                                         REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             TITLE
                                                                                                                                                                                                                                                                                       FEATURES
                                                                                                                                                                                                                              TITLE
```

Locus

용 8

占

ö

Gaps

g

```
DNA
   Immunosuppressant target proteins
Patent: US 6127521-A 28 03-OCT-2000;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ; Score 15.8; DE
; Pred. No. 89;
0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             19 bp | Sequence 28 from patent US 6509152.
AR275546
                                                            /organism="unknown"
/wol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                   19 bp | Sequence 28 from patent US 6464974.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1. .19
/organism="unknown"
/mol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    /organism="unknown"
/mol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Location/Qualifiers
                                                                                                                                                                         3689 GAGTTGCAGCAGATGTTTA 3707
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   3689 GAGTIGCAGCAGATGTTTA 3707
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               3689 GAGTTGCAGCAGATGTTTA 3707
                                                                                                                                                                                             GAGTTTGAGCAGATGTTTA 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  GAGTTTGAGCAGATGTTTA 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             GAGTTTGAGCAGATGTTTA 19
                                                                                                                                                                                                                                                                                                                ARŽ36272
ARZ36272.1 GI:27280093
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AR275546.1 GI:29708964
                                                                                                         Query Match
Best Local Similarity 89.5%;
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match 0.4%;
Best Local Similarity 89.5%;
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match 0.4%;
Best Local Similarity 89.5%;
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                               Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Unknown.
Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Unknown.
                                                                                                                                                                                                                                                                                                                                                                Unknown.
                                                                                                                                                                                                                                                                                                                                                                                Unknown.
                                                  source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       source
                                                                                                                                                                                                                                                                                                                                               KEYWORDS
SOURCE
ORGANISM
                JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                  DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                           TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               RESULT 80
AR275546
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                            REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       AUTHORS
TITLE
                                                                                                                                                                                                                                                    RESULT 79
AR236272
                                                                                                                                                                                                                                                                                                                  ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            81
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          KEYWORDS
TITLE
                                                                                                                                                                                                                                                                                                                                 VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RESULT
                                                                                                                                                                                                                                                                                     LOCUS
                                                                                                                                                                           8
                                                                                                                                                                                                      g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         엄
                                                                                                                                                                                                                                                                                                                             /note="forward primer for Japanese flounder microsatellite sequence PolillITUF"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      PAT 26-SEP-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 PAT 14-FEB-2001
                                                                         Unpublished

(Lases 1 to 21)

(Dadinza, M.R.M., Kobayashi, K., Koretsugu, S., Hasegawa, O., Ohara, E., Ozaki, A., Sakamoto, T., Naruse, K. and Okamoto, N.

Direct Submission
Submitted (14-JUN-2002) Nobuaki Okamoto, Tokyo University of Pisheries, Department of Aquatic Biosciences; 4-5-7 Konan, Minato-ku, Tokyo 108-8477, Japan

(E-mail:nokamoto@tokyo-u-fish.ac.jp, Tel:81-3-5463-0547, Pax:81-3-5463-0552)
              Coimbra, M.R.M., Kobayashi, K., Koretsugu, S., Hasegawa, O., Ohara, E., Ozaki, A., Sakamoto, T., Naruse, K. and Okamoto, N.
A genetic linkage map of the Japanease Flounder, (Paralichthys
                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Antisense modulation of casein kinase 2-beta expression Patent: US 6440738-A 110 27-AUG-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Unclassified.
1 (bases 1 to 19)
Berlin, V., Chiu, M. Isabel., Cottarel, G. and Damagnez, V.
                                                                                                                                                                                                                                                                                                                                                                              Length 21;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 linear
                                                                                                                                                                                                                                                                                                                                                                                                             3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0; Indels
                                                                                                                                                                                                                                                                                                                                                                              DB 1;
                                                                                                                                                                                                                                                  1. .21
/organism="synthetic construct"
/mol_type="genomic DNA"
d_xaref="taxon:32630"
1. .21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AR224651
Sequence 110 from patent US 6440738.
                                                                                                                                                                                                                                                                                                                                                                           Query Match 0.4%; Score 16.2; D
Best Local Similarity 85.7%; Pred. No. 96;
Matches 18; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 28 from patent US 6127521. ARI11824
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Query Match 0.4%; Score 16; DB Best Local Similarity 100.0%; Pred. No. 93; Matches 16; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1. .20
/organism="unknown"
/mol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Location/Qualifiers
                                                                                                                                                                                                                                    Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                           2522 CCGCCTTCTCAGCACACAC 2542
                                                                                                                                                                                                                                                                                                                                                                                                                                                             21 CCTCCTGCTCAGCAGTAACAC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AR224651.1 GI:23333491
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      988 CTGCTCTAGAGGCCCT 1003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AR111824.1 GI:12828672
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    16 CTGCTCTAGAGGGCCT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (bases 1 to 20)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Unclassified.
                                                               olivaceus)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Wyatt, J.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Unknown
                                                                                                                                                                                                                                                                                                               misc_feature
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                      source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Bource
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AR224651/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ORGANISM
                                                                         JOURNAL
REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              TITLE
JOURNAL
REFERENCE
AUTHORS
                                                                                                                                       TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RESULT 77
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 78
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AR111824
LOCUS
                                                                                                                                                                                                                                    FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             FEATURES
                                             TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 g
```

```
PAT 20-DEC-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               PAT 10-APR-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Length 19;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DB 1; Length 19;
 Length 19;
                                                                                                                                                                                           linear
                                   2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           2; Indels
                                                                                                                                                                                                                                                                                                                              1 (bases 1 to 19)
Berlin, V. Chiu, M.I., Cottarel, G. and Damagnez, V. Immunosuppressant target proteins
Patent: US 6464974-A 28 15-OCT-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1 (bases 1 to 19)
Berlin, V. Chiu, M.I., Cottarel, G. and Damagnez, V. Immunosuppressant target proteins
Patent: US 6509152-A 28 21-JAN-2003;
Score 15.8; DB 1;
Pred. No. 89;
0; Mismatches 2;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Score 15.8; DB 1;
Pred. No. 89;
```

ö

```
AR099520
                                                                                                                                                                                                                                                                                                                              Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  source
                                                                                                                                                                                                                                                                                                                                                                                                                                                source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           SOURCE
ORGANISM
                                                                                                                                                                                                                                      RESULT 84
AR099520/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TITLE
JOURNAL
FEATURES
REFERENCE
AUTHORS
TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                          DEFINITION
                                                                                                                                                                                                                                                                                                                                           ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 85
AR123290
                                                                                                                                                                                                                                                                                                                                                                                AUTHORS
TITLE
                                                                                                                                                                                                                                                                                       ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                     JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                    VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
                                                                                                                                                                                                                                                                  LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             셤
                                                                                                                                                                         ò
                                                                                                                                                                                               셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        PAT 31-AUG-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          PAT 29-SEP-1999
                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                            1..19
/organism="synthetic construct"
/mol type="unassigned DNA"
/mol xzef="taxon:32630"
/note=="bounstream primer used to detect the expression of the Phytoene desaturase gene by RT-PCR"
                PAT 04-OCT-2003
                                                                                                                                Giuliano, G., Rosati, C., Dharmapuri, S., Pallara, P. and Camara, B. Recombinant plants and dna constructs
Patent: EP 1323825-A 12 02-UUL-2003,
ENEA ENTE PER L NUOVE TECNOLOGIE, L'ENERGIA E L'AMBIENTE (IT)
Biogen S.r.l. (IT)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Unknow...
Unknown...
Unknown...
Unclassified.
1 (bases 1 to 20)
1 (bases 1 to 20)
Gryaznov,S.M., Schultz,R.G. and Chen,J.-k.
Gryaznotoside.
Oligonucleotide N3'-P5' phosphoramidates: hybridization and nuclease resistance properties
Patent: US 5837835-A 8 17-NOV-1998;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.4%; Score 15.8; DB 1; Length 20;
89.5%; Pred. No. 97;
                                                                                                                                                                                                                                                                                                                                  Length 19;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          linear
                   linear
                                                                                                                                                                                                                                                                                                                                                          Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          5
                                                                                                                                                                                                                                                                                                                               ch 0.4%; Score 15.8; DB 1; Similarity 89.5%; Pred. No. 89; 17; Conservative 0; Mismatches 2;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                             DNA
                    DNA

    19
/note="Pds Downstream Primer"

                                                                                                           other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20 bp
Sequence 8 from patent US 5965720.
AR079581.1 GI:10006325
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AR058876 20 bp
Sequence 8 from patent US 5837835.
AR058876
             19 bp
Sequence 12 from Patent EP1323825.
AX795182
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  3303 TATTTTTTTTTTTT 3321
                                                                                                                                                                                                                                                                                                                                                                                      1716 TCACTCTCCAGTCTTCAGG 1734
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           rarararirrirarar 19
                                                                                                                                                                                                                                                                                                                                                                                                           1 rcrcrrrccadrcrrcadd 19
                                                         AX795182.1 GI:37515943
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AR058876.1 GI:5984453
                                                                                   synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match 0.4
Best Local Similarity 89.5
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Unknown.
Unclassified.
                                                                                                                                                                                                                                                                                                                                              Best Local Similarity
Matches 17; Conserva
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Unknown.
                                                                                                                                                                                                                                                                                                primer_bind
                                                                                                                                                                                                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 83
AR079581
LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Bource
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         DEFINITION
ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          SOURCE
ORGANISM
                                                                                                                                                                                                                      source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ORGANISM
                                                                                              ORGANISM
                                DEFINITION
                                                                                                                         REFERENCE
AUTHORS
TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AUTHORS
TITLE
                                           ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 82
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AR058876
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     FEATURES
        AX795182
LOCUS
                                                                                                                                                                                                       FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           셤
                                                                                                                                                                                                                                                                                                                                                                                                              유
                                                                                                                                                                                                                                                                                                                                                                                        ð
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          PAT 16-MAY-2001
                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                         PAT 14-FEB-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                               Unclassified.

1 (bases 1 to 20)

Bennett, C. Frank. and Vickers, T.A.

Oligonuclocide compositions and methods for the modulation of the expression of BY protein

Patent: US 6077833-A 47 20-JUN-2000;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Unknown.
Unclassified.
I (bases 1 to 20)
I (bases 1 to 20)
Gryaznov, S.M., Schultz, R.G. and Chen, J.-k.
Gryaznov, S.M., Schultz, R.G. and Chen, J.-k.
Oligonucleotide N3'. fwdarw.N5'Phosphoramidate Duplexes
Oligonucleotide N3'. fwdarw.N5'Phosphoramidate Duplexes
Patent: US 6169170-A 8 02-JAN-2001;
Location/Qualifiers
                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Score 15.8; DB 1; Length 20;
Pred. No. 97;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DB 1; Length 20;
                                                                                                                                                   Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              linear
                                                                                                                                                                                                                                                                                                                                            linear
                                                                                                                                                                                       Indels
DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.4%; Score 15.8; E
89.5%; Pred. No. 97;
tive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                              20 bp 1
Sequence 47 from patent US 6077833.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       20 bp
Sequence 8 from patent US 6169170.
AR123290
AR123290.1 GI:14108256
                                                                                                                                                   Query Match 0.4%; Score 15.8; I Best Local Similarity 89.5%; Pred. No. 97; Matches 17; Conservative 0; Mismatches
                                                                                                    /organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Location/Qualifiers

    .20
    /organism="unknown"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       3303 TATTTTTTTTTTTT 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         3766 TGGCTGGGATCCCTCCCT 3784
                                                                                                                                                                                                                         3303 TATTTTTTTTTTTT 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1 rararararrirrararar 19
                                                                                                                                                                                                                                             20 TGGCTGGCATCCCTCTCT 2
                                                                                                                                                                                                                                                                                                                                                                                                AR099520.1 GI:12809286
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Query Match
Best Local Similarity 89.5%;
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
Best Local Similarity
```

the

ö

Gapa

ö

PAT 18-SEP-2002

TITLE

FEATURES

ò g

```
BD188892 20 bp DNA linear PAT 17-JUL-2003 Oligonucleotide N3' to P5' phosphoramidate: synthesis and compound; hybridization and nuclease tolerant characteristics. BD188892 BD188892 1 GI:32998631 JP 2003012688-A/8.
                                                                                                                                                                                                                                                                                                                                                                                                             Miraglia, L.J., Mero, P., Graham, M.J., Monia, B.P. and Cowsert, L.M. Antisense modulation of human MDM2 expression
Patent: JP 2002508944-A 187 26-MAR-2002;
S Unidentified INC
OS Unidentified by P 2002508944-A/187
PD 26-MAR-2002
PP 26-MAR-1999 JP 2000538025
PR 26-MAR-1999 US 09/048810
PI LOREN J MIRAGLIA, PAMELA NERO, MARK J GRAHAM, BRETT P MONIA, LEX P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   C122N15/09, A61K48/00, A61P9/10, A61P17/06, A61P3S/00, C07H21/04//
C12Q1/68,
C12N15/00
Oligonuclectide compositions and methods for the modulation of expression of B7 protein Patent: US 6319906-A 47 20-NOV-2001; Location/Qualifiers
                                                                                                                        DB 1; Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                DB 1; Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Topology: Linear;
Antisense modulation of human MDM2 expression
Location/Qualifiers
                                                                                                                                                                                                                                                                                           linear
                                                                                                                                                      Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Indels
                                                                                                                                                                                                                                                                                   20 bp DNA lin
Antisense modulation of human MDM2 expression.
BD138261
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        organism='Unidentified'
                                                                                                                                                      ..
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             2,
                                                                                                                      0.4%; Score 15.8; I
ilarity 89.5%; Pred. No. 97;
Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0.4%; Score 15.8; I 89.5%; Pred. No. 97; tive 0; Mismatches
                                                               1. .20
/organism="unknown"
/wol_type="unassigned DNA"

    .20
    /organism="unidentified"
    /mol_type="genomic DNA"
    /db_xref="taxon:32644"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           . 50
                                                                                                                                                                                 3766 TGGCTGGGATCCCTCCCCT 3784
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           2916 AAGGAACCCATGACAAAGA 2934
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Strandedness: Single;
                                                                                                                                                                                                     20 IGGCIGGCAICCCICICI 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              BD138261.1 GI:23233206
JP 2002508944-A/187.
unidentified
unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
Best Local Similarity 89.5'
                                                                                                                                                                                                                                                                                                                                                                                                   unclassified.
                                                                                                                                      Best Local Similarity
Matches 17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 5888888
                                                                                                                        Query Match
                                                                                                                                                                                                                                                           RESULT 89
BD138261/c
LOCUS
DBFINTTON
ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 90
BD188892
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                              TITLE
JOURNAL
                               JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                  REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        FEATURES
                                                 FEATURES
    TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            COMMENT
                                                                                                                                                                                                               g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         g
                                                                                                                                                                                    ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       PAT 20-APR-2002
                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                             PAT 16-MAY-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        PAT 08-AUG-2001
                                                                                                                                                                                                                                nuclear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Unknown.
Unclassified.
1 (Dases 1 to 20)
Baker,B.F., Bennett,C.Frank., Butler,M.M. and Shanahan,W.R. Jr.
Antisense oligonucleotide modulation of tumor necrosis
factor-(.alpha.) (TWF-.alpha.) expression
Patent: US 6228642-A 380 08-MAY-2001;
                                                                                                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                               1 (bases 1 to 20)
Bell,G.I., Yamagata,K., Oda,N., Kaisaki,P.J., Furuta,H.,
Horikawa,Y. and Nenzel,S.
Mutations in the diabetes susceptibility genes hepatocyte
factor (HNF) I alpha (alpha.), HNF1.beta. and HNF4.alpha
Patent: US 6187533-A 119 13-FEB-2001;
                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                       DB 1; Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Length 20;
                                                               linear
                                                                                                                                                                                                                                                                                                                                                                                    Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DB 1;
                                                               DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         DNA
                                                       Sequence 119 from patent US 6187533. ARI29530
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ARISO304 20 bp Dr
Sequence 380 from patent US 6228642.
ARISO304.1 GI:15114895
                                                                                                                                                                                                                                                                                                                                                    0.4%; Score 15.8; E
89.5%; Pred. No. 97;
tive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0.4%; Score 15.8; C
89.5%; Pred. No. 97;
tive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AR178801 20 bp C
Sequence 47 from patent US 6319906.
AR178801. GI:20219939
                                                                                                                                                                                                                                                                                                        /organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1. .20
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Unknown.
Unclassified.
1 (bases 1 to 20)
Bennett, C. Frank. and Vickers, T.A.
                                                                                                                                                                                                                                                                         Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                TTTCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                  3068 TGACTGAACTGGGTGTTCA 3086
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1 rereascaacrerecaaa 19
                                                                                                                                                                                                                                                                                                                                                                                                                                        20 TGACTGAACTGGGAGTCCA 2
                                                                                                      AR129530.1 GI:14117427
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Conservative
                                                                                                                                                                                                                                                                                                                                                                                     Conservative
                                                                                                                                                                                                                                                                                        1. .20
                                                                                                                                                                    Unclassified.
                                                                                                                                                                                                                                                                                                                                                                    Best Local Similarity
Matches 17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
Best Local Similarity
Matches 17; Conserv
                                                                                                                                    Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                3414
                                                                                                                                                                                                                                                                                                                                                         Query Match
                                                        LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                            source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AR150304
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RESULT 88
AR178801/c
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           source
                              RESULT 86
AR129530/c
                                                                                                                                                   ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            REFERENCE
                                                                                                                                                                                 REFERENCE
                                                                                                                                                                                                 AUTHORS
                                                                                                                                                                                                                                                           JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT 87
```

FEATURES

ઠ g

ö

Gaps

ö

Key

E

```
PAT 06-FEB-1997
                                                                                                                                                                                                                                                                                                                                                                                                   PAT 24-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1 (bases 1 to 20)
Gryaznov, S.M., Schultz, R.G. and Chen, J.-k.
Gligonuclectide N3. Fwdarw.P5' phosphoramidates: triplex DNA formation
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           .
0
                                                                                                                                                                                                   .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match 0.4%; Score 15.8; DB 1; Length 20; Best Local Similarity 89.5%; Pred. No. 97; Matches 17; Conservative 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Length 20;
/organism='Artificial Sequence'
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                linear
                                                                                                                                                                                                                                                                                                                                                                                                     linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Indels
                                                                                                                                                                                                   Indels
                                                                                                                                                                                                   2;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               5,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DB 1;
                                                                                                                                                            DB 1;
                                                   1. .20
/organism="synthetic construct"
/mol_type="genomic DNA"
/db_xref="taxon:32630"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Diagnostic test
Patent: WO 2004018711-A 34 04-MAR-2004;
University College London (GB)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                DNA
                                                                                                                                                                                                                                                                                                                                                                                              CQ786110 20 bp DNJ Sequence 34 from Patent WO2004018711.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          other sequences; artificial sequences
                                                                                                                                                       ; Score 15.8; DE; Pred. No. 97; 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match 0.4%; Score 15.8; Dest Local Similarity 89.5%; Pred. No. 97; Matches 17; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1. .20
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Patent: US 5591607-A 8 07-JAN-1997;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       20 bp
Sequence 8 from patent US 5591607.
133253
                              Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     3303 TATTTTATTTTATATA 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1995 ACACCTTCAGATAAGCAGG 2013
                                                                                                                                                                                                                                                3414 TITCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                            rcrcaaccaacrcrccaaa 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20 Acacciccadaaaadcadd 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      CQ786110.1 GI:45721213
                                                                                                                                                         Ouery Match
Best Local Similarity 89.5%;
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  I33253.1 GI:1824044
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Ming-Qing, D.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Unknown
         H
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              source
                                                                                                                                                                                                                                                                                                                                                                                                                            DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                       source
                                                                                                                                                                                                                                                                                                                                                                RESULT 92
CQ786110/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 93
133253
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    FEATURES
                                FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ð
                                                                                                                                                                                                                                                         ò
                                                                                                                                                                                                                                                                                            g
    unclassified
unclassified.
1 (bases 1 to 20)
SG Tyaznov, S.M., Schultz, R.G. and Chen, J.
SGryaznov, S.M., Schultz, R.G. and Chen, J.
Oligonucleotide N3' to P5' phosphoramidate: synthesis and compound hybridization and nuclease tolerant characteristics
Patent: JP 2003012688-A 8 15-JAN-2003;
LYNX THERAPEUTICS INC
OS Unidentified
PN JP 2003012688-A/8
PD 15-JAN-2003
PF 12-JUN-2002
PP 2003012688-A/8
PR 18-MAR-1994 US 08/210505, 18-MAR-1994 US 08/214599 PI SERGEI M GRYAZNOV, RONALD G SCHULTZ, JER-KANG CHEN PC
COTH99/15/(1201/02, 1201/68)
CC Topology: Linear;
CC Topology: Lin
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      BD22817.1 GI:33037947
JP 2002526125-A/380.
Synthetic construct
synthetic construct
construct
Other sequences; artificial sequences.
E 1 (bases 1 to 20)
Baker, B.F., Bennett, F.C., Butler, M.M. and Jr, W.J.S.
Antisense oligonuclectide regulation of expression of tumor necrosis factor-alpha (TNF-alpha)
L. Patent: JP 2002526125-A 380 20-AUG-2002;
ISIS PHARMACEUTICALS INC
OS Artificial Sequence
PN JP 2002526125-A/380
PD 20-AUG-2002
PP 05-OCT-1999 JP 2000574737
PR 05-OCT-1999 US 09/166186,18-MAY-1999 US 09/313932 PI
BRENNA F BAKER, FRANK C BENNETT, MADELINE M BUTLER, WILLIAM J PI
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Antisense oligonucleotide regulation of expression of tumor necrosis factor-alpha (TNF-alpha).
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          CINIS/09, AGIK31/7115, AGIK31/712, AGIK31/7125, AGIK48/00, AGIP1/
00, AGIP1/16,
AGIP1/18, AGIP3/10, AGIP17/00, AGIP17/04, AGIP29/00, AGIP31/00, PC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            5
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Score 15.8; DB 1;
Pred. No. 97;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1. .20
/organism="unidentified"
/mol_type="genomic DNA"
/db_xref="taxon:32644"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             reature 7. .8.
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              SHANAHAN JR
PC C12N15/09,A61K31/7115,
PC 00,A61P1/16,
PC A61P1/16,
C07H21/02,
PC C07H21/04,C12N15/00
CC Synthetic Locat:
FH RESULT FH SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         3303 TATTTTTTTTTTAT 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ch 0.4%;
1. Similarity 89.5%;
17; Conservative 0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match
Best Local Similarity
Matches 17; Conserva
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 91
BD228177
                    ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      JOURNAL
                                                            REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
                                                                                                                                                  JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  COMMENT
                                                                                                           TITLE
                                                                                                                                                                                               COMMENT
```

셤

ö

```
Anctil, J.L. and Cote, G. Molecular diagnostic of glaucomas associated with chromosomes 2 and Molecular diagnostic of glaucomas
                                                                                                                                       ö
                                                                                                                                                                                                                                                                            PAT 18-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         PAT 24-AUG-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                       Gape
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Gape
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                           DB 1; Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DB 1; Length 20;
                                                                                                                                                                                                                                                                             linear
Oligodeoxyribonucleotide N3' P5' phosphoramidates
Patent: US 5726297-A 8 10-MAR-1998;
Location/Qualifiers
                                                                                                                                       2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         linear
                                                                                                                                                                                                                                                                                                                                                                                               1 (bases 1 to 20)
Dean, N.M., Marcusson, B.G. and Wyatt, J.
Antisense modulation of Fas mediated signaling
Patent: US 6653133-A 153 25-NOV-2003;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1. .20
/organism="synthetic construct"
/mol_type="unassigned DNA"
/db xref="taxon:32610"
/noEe="OLIGONUCLEOTIDE"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Patent: WO 9916899-A 22 08-APR-1999;
ANCTIL JEAN LOUIS (CA); COTE GILLES (CA)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              synthetic construct
synthetic construct
other sequences; artificial sequences.
                                                                                                                                                                                                                                                                    20 bp Di
Sequence 153 from patent US 6653133.
AR432353.1 GI:40194626
                                                                                                        0.4%; Score 15.8; I
al Similarity 89.5%; Pred. No. 97;
17; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match

0.4%; Score 15.8; E
Best Local Similarity 89.5%; Pred. No. 97;
Matches 17; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match 0.4%; Score 15.8; E
Best Local Similarity 89.5%; Pred. No. 97;
Matches 17; Conservative 0; Mismatches
                                               1. .20
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20 bp
Sequence 22 from Patent W09916899.
AX004440.
AX004440.1 GI:9927899
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1. .20
/organism="unknown"
/mol_type="genomic DNA"
                                                                                                                                                                    3303 TATTITITITITITI 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        3735 GGACCTATGAATCTATTTA 3753
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 861 TTAAGAAATAATTTTGATA 879
                                                                                                                                                                                     1 TATATATTTTTATATAT 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       20 TTAAGAAATAATATTATA 2
                                                                                                                                                                                                                                                                                                                                                                                    Unclassified.
                                                                                                                         Best Local Similarity
Matches 17; Conserv
                                                                                                                                                                                                                                                                                                                                                      Unknown
                                                                                                                                                                                                                                                                                                                                                                      Unknown
                                                                                                           Query Match
                                                                                                                                                                                                                                                                                                                                      KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DEFINITION
ACCESSION
VERSION
                                                                                                                                                                                                                                              RESULT 97
AR432353/c
 TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ORGANISM
                                                                                                                                                                                                                                                                                          DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
AUTHORS
TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                        ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RESULT 98
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AX004440
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   SOURCE
                                                                                                                                                                                                                                                                            LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ઠે
                                                                             PAT 13-MAY-1997
                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Unclassified.
Unclassified.
1 (bases 1 to 20)
Gryaznov.S.M., Schultz,R.G. and Chen,J.-K.
Gryaznov.S.M., Schultz,R.G. and Chen,J.-K.
Oligonucleotide N3..fwdarw.P5' phosphoramidates: hybridization and nuclease resistance properties
Patent: US 5631135-A 8 20-MAY-1997;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        PAT 07-0CT-1997
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    PAT 01-DEC-1998
                                                                                                                                                                                                                                                                                                                                                                                    Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                           Unclassified.

1 (bases 1 to 20)
Gryaznov, S.M., Schultz, R.G. and Chen, J.-k.
Gryaznov, S.M., Schultz, R.G. and Chen, J.-k.
Oligonucleotide Na3-P5' phosphoramidates: hybridization and nuclease resistance properties
Patent: US 5599922-A 8 04-PEB-1997;
                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                      Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Length 20;
                                                                             linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  linear
                                                                                                                                                                                                                                                                                                                                                                                  Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Unclassified.
1 (bases 1 to 20)
Gryaznov,S.M., Schultz,R.G. and Chen,J.-K.
                                                                                                                                                                                                                                                                                                                                                   Score 15.8; DB 1;
Pred. No. 97;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                DB 1;
                                                                             DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Ouery Match 0.4%; Score 15.8; D
Best Local Similarity 89.5%; Pred. No. 97;
Matches 17; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                        1. .20
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1. .20
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20 bp
Sequence 8 from patent US 5726297.
192006
                                                                        20 bp
Sequence 8 from patent US 5599922.
135518
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       143128 20 bp
Sequence 8 from patent US 5631135.
                                                                                                                                                                                                                                                                          Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                               3303 TATTTTTTTTTTTT 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          3303 TATTTTTTTTTTATAT 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                1 TATATATTTTTATATAT 19
 TATATATTTTTATAT 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%;
                                                                                                                      I35518.1 GI:2088486
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   GI:2468372
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                GI:3936476
                                                                                                                                                                                                                                                                                                                                                 Query Match
Best Local Similarity 89.57
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   143128.1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                192006.1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Unknown.
                                                                                                                                                  Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Unknown
                                                                                                                                                                     Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Unknown
                                                                                                    ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                           I35518
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                          source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ORGANISM
                                                                                                                                                                                                              AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 96
192006
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AUTHORS
TITLE
                                                                                                                                                                                                                                                       JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       JOURNAL
                                                                                                                                                                                                REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 95
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             REFERENCE
                                             RESULT 94
                                                                                                                                                                                                                                                                          FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       FEATURES
```

a

셤

셤

```
PAT 07-SEP-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Oligonucleotide for amplification and detection of Epstein-Bar
                                                                                                                                                                                                                                                                                                                                                                                                      PAT 18-SEP-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             98204231.9 PI
ADRIANUS
PI MIDDELDORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Human herpesvirus 4 (Epstein-Barr virus)
Human herpesvirus 4
Viruses; dsDNA viruses, no RNA stage; Herpesviridae;
Gammaherpesvirinae; Lymphocryptovirus.
1 (bases 1 to 21)
Vervoort, M.B.H.J. Den, A.J.C.V. and Middeldorp, J.M.
Oligomucleoride for amplification and detection of Epstein-Bar
virus (EBV) nucleic acid
Patent: JP 2002505122-A 25 19-FEB-2002;
AKZO NOBEL NV
                                                                                                                                                                                                                                                                                                                                                                                                    BD136927 21 bp DNA linear PAT 18-SEP-Oligonucleotide for amplification and detection of Epstein-Bar virus (EBV) nucleic acid.
BD136927
                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ;
0
                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /organism='Epstein-barr virus'.
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
0.4%; Score 15.8; DB 1; Length 21;
Best Local Similarity 89.5%; Pred. No. 1e+02;
Matches 17; Conservative 0; Mismatches 2; Indels
                                                                                                                                                                                       Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     linear
                                                                                                                                                                                                                                  Indels
                                                                                                                                                                                                                                  2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /organism="Human herpesvirus 4"
/mol_type="genomic DNA"
/db_xref="taxon:10376"
                                                                                                                                                                                           1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Location/Qualifiers
                                                      Location/Qualifiers.
                                                                                                                                                                                           B
                                                                                                                                                                                         ch 0.4%; Score 15.8; I Similarity 89.5%; Pred. No. 97; 17; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AX018466 25 from Patent WO9945155.

    .20
    /organism="Homo sapiens"
/mol_type="genomic DNA"
/db_xref="taxon:9606"

                                                                           Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   . .21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2319 GCTGCCAACATCATCCATG 2337
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            virus (EBV)
                                                                                                                                                                                                                                                                        516 GCCCTGGGAGCCTTTCTG 534
                                                                                                                                                                                                                                                                                                   1 GCCCTGGTAGCCTTCTCTG 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             21 GCTGCCAACATCTCCCATG 3
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        BD136927.1 GI:23231872
                A61P35/00,
synthesized primer
CC A61P35
CC sequence
FH Key
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              nucleic acid
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     source
                                                                                                                                                                                                                 Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            SSEEE
                                                                                                                                                                                                 Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AX018466/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DEFINITION
                                                                                                                                                                                                                                                                                                                                                                        RESULT 101
BD136927/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RESULT 102
                                                                                                                                                                                                                                                                                                                                                                                                                 LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                      Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            FEATURES
                                                                                 FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            COMMENT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         8
                                                                                                                                                                                                                                                                                                               g
                                                                                                                                                                                                                                                                              8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Eutheleostomi; Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.

E. 1 (Dases 1 to 20)
S. Ota, T., Isogai, T., Nishikawa, T., Hayashi, K., Saito, K., Yamamoto, J., Ishii, S., Sugiyama, T., Makamatsu, A., Nagai, K., Otsuki, T., Kishimoto, T., Yano, K., Kanzaki, K. and Inoue, Y. Kishimoto, T., Yano, K., Kanzaki, K. and Inoue, Y. Guanosine triphosphate-binding protein-coupled receptors, genes thereof and production and use of the same.

I. Patent: WO 109323-A 20 08-FEB-2001; HELIX RESEARCH INSTITUTE, TOSHIO OTA, TAKAO ISOGAI, TETSUO NISHIKAWA, KOJI HAXASHI, KAORU SAITO, UNICHI YAMAMOTO, SHIZUKO ISHII, OMOYASU SUGIYAMA, AI WAKAMATSU, KEIICHI NAGAI, TETSUJI OTSUKI, TOSHIMITSU NI WO 0109323-A.O.

PD 08-FEB-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       TOSHIO
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                PAT 02-AUG-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    PF 28-JUL-2000 WO 2000JP005070
PR 29-JUL-2009 JP 99P 248036,27-AUG-1999 JP 99P 300253 PR
11-JAN-2000 JP 00P 118776,02-MAY-2000 JP 00P 183767 PR
18-OCT-1999 US 60/159590,17-FEB-2000 US 60/183322 PI TOSHICO OTA,TAKADO ISOGAI, TETSUO NISHIKAMA, KOJI HAYASHI, PI KAORU SAITO, PI JUNICHI YAMAMOTO,SHIZUKO ISHI, TOOMOYASU SUGIYAMA, AI WAKAWATSU, PI KEIICHI NAGAI,TETSUJI OTSUKI,TOSHIMITSU KISHIMOTO, PI KAZUHIRO YANO,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       KOJI KANŽAKI, YOSHIHISA INOUB
CLDANIS/12, CL2NIS/63, CL2P21/02, CO7K14/705, CO7K16/28, A61K45/00,
A61P25/00,
A61P25/28, GOIN33/566, GOIN33/50, GOIN33/15
Description of Artificial Sequence: an artificially
                                                                                                     PAT 26-MAR-2003
                                                                                                                                                                                                                                                                              Okuda, T., Saito, S., Dorsey, K.M. and Tsuzaki, Y. Modified dna molecule, recombinant containing the same thing, and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        BD012503 20 bp DNA linear PAT 02-AUG-:
Guanosine triphosphate-binding protein-coupled receptors, genes
thereof and production and use of the same.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Length 20;
                                                                                                         linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         7
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                   /organism="synthetic construct"
/mol type="unassigned DNA"
/mol type="unassigned DNA"
/note="primer M11-7"
                                                                                                         DNA
                                                                                                                                                                                                                                          other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                         uses thereof
Patent: EP 1275716-A 36 15-JAN-2003;
Zeon Corporation (JP)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ch 0.4%; Score 15.8; I Similarity 89.5%; Pred. No. 97; 17; Conservative 0; Mismatches
                                                                                                 20 bp
Sequence 36 from Patent BP1275716.
AX665199
                                                                                                                                                                                                                                                                                                                                                                              Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              3702 IGTTTATTTTTCAGAAC 3720
            GGAACTGTGAATCTATTTA 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           BD012503.1 GI:22092692
WO 0109323-A/20.
Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  19 rerrratrererreadade 1
                                                                                                                                                                AX665199.1 GI:29290323
                                                                                                                                                                                                         synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           BD012503
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 100
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ORGANISM
                                                                                                                        DEFINITION
                                                                                       AX665199/c
                                                                                                                                                ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                           ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                      AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ACCESSION
                                                                                                                                                                                                                                                                  REFERENCE
                                                                                                                                                                                                                                                                                                                                             JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                VERSION
KEYWORDS
                                                                       RESULT 99
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    BD012503
                                                                                                                                                                                                                                                                                                                                                                                    PEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 COMMENT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           TITLE
                                                                                                                                                                                                           SOURCE
```

셤 ð

ö

Gaps

Gaps

ö

PAT 07-0CT-1997

ACCESSION

REFERENCE AUTHORS

TITLE

JOURNAL

FEATURES

```
Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Telerman, A., Amson, R. and Tuijnder, M. Sequences involved in phenomena of tumour suppression, tumour reversion, apoptosis and/or virus resistance and their use as
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Unclassified.

1 (bases 1 to 17)

Stinchcomb,D.T. Draper,K., McSwiggen,J. and Jarvis,T.
C-myb targeted ribozymes
Patent: US 5646042-A 1048 08-JUL-1997;
Location/Qualifiers
                                                                                                                   1; Length 17;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DB 1; Length 17;
                                                                                                                                                         1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Patent: WO 03025175-A 1236 27-MAR-2003;
Molecular Engines Laboratories (FR)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AX729602 17 bp DNA Sequence 1236 from Patent W003025175. AX729602
                                                                                                                                                                                                                                                                                                                                 153307 17 bp DNA Sequence 1048 from patent US 5646042.
JOURNAL Patent: US 5817796-A 1048 06-OCT-1998;
SATURES Location/Qualifiers
                                                                                                                   四
                                                                                                                   0.4%; Score 15.4; I 94.1%; Pred. No. 82; tive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%; Score 15.4; D
94.1%; Pred. No. 82;
ive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match 0.4%; Score 15.4; D
Best Local Similarity 94.1%; Pred. No. 82;
Matches 16; Conservative 0; Mismatches

    .17
    /organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"

                                          1. .17
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                              3305 TTTTTATTTTATAT 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  3305 TTTTTATTTTATAT 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  2411 TGAATATGAGATTGCTC 2427
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AX729602.1 GI:30508945
                                                                                                                                                                                                                                  1 ririraaririaarar 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1 Tritraarritarar 17
                                                                                                                                                                                                                                                                                                                                                                                            153307.1 GI:2474510
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens (human)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Ouery Match
Best Local Similarity 94.15
Matches 16; Conservative
                                                                                                               Query Match
Best Local Similarity 94.1
Matches 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               medicines
                                                                                                                                                                                                                                                                                                                                                                                                                                     Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                         Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                   SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             source
                                                                                                                                                                                                                                                                                             RESULT 105
I53307
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 106
AX729602/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                      DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AUTHORS
TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                        ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           REFERENCE
                      FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    LOCUS
                                                                                                                                                                                                                                                                                                                                      LOCUS
                                                                                                                                                                                                                                  g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ð
                                                                                                                                                                                                Š
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /organism="synthetic construct"
Mol type="unassigned DNA"
/db xref="trans12650"
/note="Description of Artificial Sequence: oligonucleotide
                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          PAT 29-SEP-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             PAT 18-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                               Middeldorp, J.M., Van Den Brule, A.J. and Vervoort, M.B. Oligonucleotides for the amplification and detection of epstein obarr virus (ebv) nucleic acid Patent: WO 9945155-A 25 10-8EP-1999; MIDBELDORP JAAP MICHIEL (NL); AKZO NOBEL NV (NL); DEN BRULE ADRIANUS JOHANNES CH (NL); VERVOORT MARCEL BARTOLINA HEND (NL) Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Unclassified.
1 (Dases 1 to 17)
Stinchcomb.D.T., Draper.K., McSwiggen,J. and Jarvis,T.
C-myb ribozymes having 2'-5'-linked adenylate residues
                                                                          Human herpesvirus 4
Viruses; dsDNA viruses, no RNA stage; Herpesviridae;
Gammaherpesvirinae; Lymphocryptovirus.
                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15.8; DB 1; Length 21; 89.5%; Pred. No. 1e+02; tive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%; Score 15.8; DB 1; Length 21; 89.5%; Pred. No. 1e+02; tive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Nov-x proteins and nucleic acids encoding same Patent: WO 02068652-A 476 06-SEP-2002; Location/Qualifiers
                                                                                                                                                                                                                                                                                           1. .21
/organism="Human herpesvirus 4"
/mol_type="unassigned DNA"
/db_xref="taxon:10376"
                                                        Human herpesvirus 4 (Epstein-Barr virus)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 476 from Patent W002068652.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AR046255
Sequence 1048 from patent US 5817796.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2319 GCTGCCAACATCATCCATG 2337
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  2626 GGAATCCAGAAGGAACAGT 2644
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             21 GCTGCCAACATCTCCCATG 3
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               21 GGAAACCAAGGAACAGT 3
  AX018466
AX018466.1 GI:10042617
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AX921483.1 GI:40215104
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AR046255
AR046255.1 GI:5967720
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                             17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Best Local Similarity 89.5
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                  Query Match
Best Local Similarity
Matches 17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
                    VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AR046255
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AX921483/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DEFINITION
```

ACCESSION VERSION KEYWORDS

RESULT 103

ઠે g ORGANISM

REFERENCE AUTHORS JOURNAL

TITLE

PEATURES

ö

Gарв

ö

PAT 08-MAY-2003

ö

Gaps

ö

ACCESSION VERSION KEYWORDS SOURCE ORGANISM

RESULT 104

엄

REFERENCE AUTHORS TITLE

g

```
unidentified
unidentified
unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Best Local Similarity
Matches 16; Conserva
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       purposes
                                                                                                                                                                                                                                                        AX637740
                                                                                                                                                                                                                                                                                                                                                                                                                           genes
                                                                                                                                                            N
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                RESULT 110
AX637740/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   source
                                                                                                                                                                                                                                         DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                             SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                CQ788034/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 111
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ACCESSION
VERSION
        JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                            JOURNAL
                                                                                                                                                                                                                                                                                                                                     REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     PEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ò
                                                                                                                                       ð
                                                                                                                                                            a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     01-SEP-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                PAT 01-SEP-2000
                                                                                                                                                                   l (bases 1 to 18)
Sullivan, S., Draper, K., Kisich, K., Stinchcomb, D.T. and McSwiggen, J.
TNF-.alpha. ribozymes
Patent: US 5811300-A 1081 22-SEP-1998;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                              ö
                                                             PAT 29-SEP-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Unknown.
Unclassified.
I (bases 1 to 18)
Bohinski,R.J. and Whitsett,J.A.
Nucleic acid sequences controlling lung cell-specific gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Unknown.
Unclassified.
1 (bases 1 to 18)
Bohinski, R.J. and Whitsett, J.A.
Nucleic acid sequences controlling lung cell-specific gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     PAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match 0.4%; Score 15.4; DB 1; Length 18; Best Local Similarity 94.1%; Pred. No. 89; Matches 16; Conservative 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                    Query Match
0.4%; Score 15.4; DB 1; Length 18;
Best Local Similarity 94.1%; Pred. No. 89;
Matches 16; Conservative 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     linear
                                                             linear
                                                                                                                                                                                                                                                                                                                                                                                                                                    linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                    DNA
                                                           AR042291 18 bp DNA Sequence 1081 from patent US 5811300.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             expression
Patent: US 5976873-A 56 02-NOV-1999;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AR083519 18 bp D
Sequence 58 from patent US 5976873.
AR083519
AR083519.1 GI:10010292
                                                                                                                                                                                                                                                                                                                                                                                                                              18 bp 1
Sequence 56 from patent US 5976873.
AR083517
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1. .18
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                    1. .18
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1798 CCCTCCAGGTTCTTGAT 1814
                                                                                                                                                                                                                                                                                                                                          3416 TCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AR083517.1 GI:10010290
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             CCCTCCAGGTGCTTGAT 1
                                                                                                   AR042291.1 GI:5962787
                                                                                                                                                                                                                                                                                                                                                                   18 rchaddaadrcrddaaa 2
17 TGAATATGAGATTGATC 1
                                                                                                                                                        Unclassified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Unknown.
                                                                                                                                 Unknown.
                                                                                                                                            Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                    LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 109
AR083519
LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                             LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                      source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     SOURCE
ORGANISM
                                                                                                                                                                                              TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                            RESULT 108
AR083517/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          REFERENCE
AUTHORS
TITLE
                                                    AR042291/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               JOURNAL
                                      RESULT 107
                                                                                                                                                                      REFERENCE
                                                                                                                                                                                  AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            FEATURES
                                                                                                                                                                                                                         FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      g
                                                                                                                                                                                                                                                                                                                                                                     g
```

à

```
Stinchcomb, D.T., Dudycz, L.W., Chowrira, B., Grimm, S., Direnzo, A., Karpeisky, A., Draper, K.G., Kisich, K., Matulic-Adamic, J., Mcswiggen, J.A., Modak, A., Pavco, P., Beigelman, L., Sullivan, S.M., Sweiger, D., Thompson, J.D., Tracz, D., Usman, N., Wincott, F.E. and Woolf, T.
Method and reagent for inhibiting the expression of disease related
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PAT 24-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                 PAT 21-FEB-2003
                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Geldermann,H., Preuss,S. and Han,Y.
Polymorphous microsatellite loci in genes for pre-diagnostic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                  .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DB 1; Length 18;
                                                                                                                                                         Length 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Indels
                                                                                                                                                                                                                                                                                                                                                                                                         linear
                                                                                                                                                                                                  1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ä
                                                                                                                                                     Query Match
0.4%; Score 15.4; DB 1;
Best Local Similarity 94.1%; Pred. No. 89;
Matches 16; Conservative 0; Mismatches 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          purposes
Patent: WO 2004020664-A 340 11-MAR-2004;
Universitaet Hohenheim (DE)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 CQ788034 19 bp DNA Sequence 340 from Patent WO2004020664.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Patent: EP 1260586-A 4879 27-NOV-2002;
RIBOZYME PHARMACEUTICALS, INC. (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                      AX637740 18 bp R Sequence 4879 from Patent EP1260586.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 1. .19
/organism="Homo sapiens"
/mol_type="unassigned DNA"
/db_xref="taxon:9606"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ch
1 Similarity 94.1%; Pred. No. 89;
16; Conservative 0; Mismatches
expression
Patent: US 5976873-A 58 02-NOV-1999;
Location/Qualifiers

    .18
    /organism="unidentified"
    /mol_type="unassigned RNA"
    /db_xref="taxon:32644"

                                                                    1. .18
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           3416 TCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                   1798 CCCTCCAGGTTCTTGAT 1814
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  CQ788034
CQ788034.1 GI:45722987
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AX637740.1 GI:28473354
                                                                                                                                                                                                                                                                           CCCTCCAGGTGCTTGAT 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens (human)
```

PAT 17-JUL-2003

```
other sequences; artificial sequences.

E 1 (bases 1 to 20)
S Baker, Ber. F., Bendert, F.C., Butler, M.M. and Jr, W.J.S.
Antisense oligonucleotide regulation of expression of tumor necrosis factor-alpha (TNF-alpha)
L PSIS PHARMACEUTICALS INC
OS Artificial Sequence
PN JP 2005256125-A/228
PN JP 2005256125-A/228
PN JP 200526125-A/228
PN JP 200556125-A/228
PN JP 200556125-A/2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           C12N15/09, A61K31/7115, A61K31/712, A61K31/7125, A61K48/00, A61P1/
00, A61P1/16,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             A61P1/18, A61P3/10, A61P17/00, A61P17/04, A61P29/00, A61P31/00, PC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Synthetic construct
basea: 1 to 20
construct
c
                                                                                                                                                                                                                                                                                        Antisense oligonucleotide regulation of expression of tumor necrosis factor-alpha (TNF-alpha).
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      BD235863 20 a linear PAT 1:
Method of estimating bone fracture frequency by screening
polymorphism in vitamin D receptor gene.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /organism='Artificial Sequence'
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.4%; Score 15.4; DB 1; Length 20; 94.1%; Pred. No. 1e+02; ive 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /organism="synthetic construct"
/mol_type="genomic DNA"
/db_xref="taxon:32630"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        C07H21/04,C12N15/00
   3416 TCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          3416 TCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                               BD228025.1 GI:33037795
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  BD235863.1 GI:33045633
JP 2002525074-A/4.
                                            1 TCAAGGAAGTCTGGAAA 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   l Similarity 94.1%;
16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                    JP 2002526125-A/228.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Best Local S
Matches 16
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     BD235863/c
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                      ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        SOURCE
ORGANISM
                                                                                                                                                                             RESULT 114
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 115
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  VERSION
KEYWORDS
                                                                                                                                                                                                                  BD228025
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           COMMENT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  COMMENT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           PAT 14-FEB-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  PAT 08-AUG-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1 (bases I to 20)
Baker, B.F., Bennett, C.Frank., Butler, M.M. and Shanahan, W.R. Jr. Antisense oligonucleotide modulation of tumor necrosis factor-alpha. (TNF-alpha.) expression
Patent: US 600580-A 228 27-JUN-2000;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1 (bases 1 to 20)
Baker, B.F., Bennett, C. Frank., Butler, M.M. and Shanahan, W.R. Jr.
Antisense oligonucleotide modulation of tumor necrosis
factor-(.alpha.) (TNF-.alpha.) expression
Patent: US 6228642-A 228 08-MAY-2001;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gapa
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match
0.4%; Score 15.4; DB 1; Length 20;
Best Local Similarity 94.1%; Pred. No. 1e+02;
Matches 16; Conservative 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match 0.4%; Score 15.4; DB 1; Length 20; Best Local Similarity 94.1%; Pred. No. 1e+02; Matches 16; Conservative 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                               Length 19;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1; Indels
                                                                             1. .4
/note="Anzahl der Wiederholungen: 4"
                                                                                                                                            9. .12
/note="Anzahl der Wiederholungen: 5"
                                                                                                                                                                                                                  13. .15
/note="Anzahl der Wiederholungen: 1"
                                                                                                                                                                                                                                                                                    16. .19
/note="Anzahl der Wiederholungen: 1"
                                                                                                                                                                                                                                                                                                                                                                                        Query Match
0.4%; Score 15.4; DB 1;
Best Local Similarity 94.1%; Pred. No. 97;
Matches 16; Conservative 0; Mismatches 1;

    .19
/note="M09, Allel R (PrP-Gen)"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20 bp Di
Sequence 228 from patent US 6228642.
AR150152
AR150152.1 GI:15114743
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 228 from patent US 6080580. AR100497
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       /organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       /mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1. .20
/organism="unknown"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1061 TATTATTGACTTATTA 1077
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              3416 TCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AR100497.1 GI:12810945
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 rcaaggaagrcrggaaa 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        17 TATTATTACTTATTA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Inclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Unknown,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Unknown
                                                                         repeat_unit
                                                                                                                                            repeat_unit
                                                                                                                                                                                                                  repeat_unit
                                                                                                                                                                                                                                                                                repeat_unit
satellite
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        DEFINITION
ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 112
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RESULT 113
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           JOURNAL
```

REPERENCE

FEATURES

AR100497

ሯ 윱 LOCUS

AR150152 LOCUS

ò g REFERENCE

ö

Gaps

ö

PAT 17-JUL-2003

Gaps

.. 0

PAT 07-0CT-1996

슝

```
Including the 20)
Gallie, B.L., Dunn, J.M. and Stevens, J.K.
Method, reagents and kit for diagnosis and targeted screening for retinoblastoma
retinoblastoma
Patent: US 5550020-A 100 27-AUG-1996;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Unclassified.

1 (bases 1 to 20)

Whitcomb, D., Ehrlich, G.D. and Gorry, M.C.
Method for determining whether a human patient is susceptible to hereditary pancreatitis, and primers therefore
Patent: US 6406846-A 41 18-JUN-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match 0.4%; Score 15.4; DB 1; Length 20; Best Local Similarity 94.1%; Pred. No. 1e+02; Matches 16; Conservative 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match 0.4%; Score 15.4; DB 1; Length 20; Best Local Similarity 94.1%; Pred. No. 1e+02; Matches 16; Conservative 0; Mismatches 1; Indels
                                                                                                                                                                        Length 20;
                                                                                                                                                                                                                                                                                                                                                                   linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             linear
                                                                                                                                                                                                           1; Indels
                                                                                                                                                                      Query Match

0.4%; Score 15.4; DB 1;
Best Local Similarity 94.1%; Pred. No. 1e+02;
Matches 16; Conservative 0; Mismatches 1;
                                                                                  /organism="synthetic construct"
/mol type="unasaigned DNA"
/mol type="unasaigned"
/mole="reverse primer"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DNA
iii)
Patent: WO 2004074830-A 4 02-SEP-2004;
Payer HealthCare AG (DE)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                           20 bp Di
Sequence 100 from patent US 5550020.
[25313
[25313.1 GI:1605183
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      20 bp 1
Sequence 41 from patent US 6406846.
AR213949
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1. .20
/organism="unknown"
/mol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1. .20
/organism="unknown"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1830 ATAATGACTTCCAAAAA 1846
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1098 TCATTTTCCTGGTGAG 1114
                                                                                                                                                                                                                                               2834 TCAAGGAGCTTCCAGTG 2850
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AR213949.1 GI:23311368
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            4 ATAATAACTTCCAAAAA 20
                                                                                                                                                                                                                                                                              1 rcardeadcriccadrd 17
                                                                       1. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                            Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 119
AR213949/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       SOURCE
ORGANISM
                                                                           source
                                                                                                                                                                                                                                                                                                                                     RESULT 118
125313
                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                     DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                         ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   JOURNAL
                     JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             FEATURES
                                                        FEATURES
                                                                                                                                                                                                                                                                                                                                                                          LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           셤
                                                                                                                                                                                                                                                                                  셤
                                                                                                                                                                                                                                                     à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Golz,S., Brueggemeier,U. and Summer,H.
Diagnostics and therapeutics for diseases associated with g-protein
coupled receptor prostaglandin e2 ep3 iii (prostaglandin e2 ep3
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               PAT 13-SEP-2004
   10-SEP-1999 JP 2000570366
10-SEP-1998 GB 9819769.2
ANDREAS GERARDUS UITTERLINDEN, JOHANNES PETRUS THOMAS MARIA VAN
                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               PAT 05-JUL-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Schwenzer, B., Schmidt, U., Wirth, M.P., Kraemer, K., Fuessel, S. and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ;
0
                                                                                                                                                                                                                                                                                                                                           .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Meye,A.
Polymicleotides targeted against htert and use thereof
Polymicleotides targeted 3 24-UIN-2004;
Petent: WO 2004053116-A 5 24-UIN-2004;
Technische Universitaet Dresden (DE)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 1. .20
/organism="synthetic construct"
/mol type="unassigned DNA"
/db xref="taxon:32630"
/note="Beschrelbung der k nstlichen Sequenz:
anti-hTERT-AS-Konstrukt"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%; Score 15.4; DB 1; Length 20;
94.1%; Pred. No. 1e+02;
tive 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                    Query Match
0.4%; Score 15.4; DB 1; Length 20;
Best Local Similarity 94.1%; Pred. No. 1e+02;
Matches 16; Conservative 0; Mismatches 1; Indels
                                                                                                                                                                               /organism='Artificial Sequence'
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   linear
                                                                                         HUIBERT ADRIAAN PIETER POLS
C12N15/09,C12Q1/68,C12N15/00
Description of Artificial Sequence:Synthetic
                                                                                                                                                                                                                      1. .20
/organism="synthetic construct"
/mol type="genomic DNA"
/db_xref="taxon:32630"
                                                                                                                                                Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     synthetic construct
synthetic construct
other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     synthetic construct
synthetic construct
other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20 bp DP Sequence 4 from Patent WO2004074830. CQ867622 GI:51997814
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 CQ829824 20 bp DN
Sequence 5 from Patent W02004053116.
CQ829824
CQ829824.1 GI:49732949
                                                                                                                                                                                                                                                                                                                                                                              2209 GCCAGGATGAGGACTGG 2225
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            468 AGCCACCTCTCTACCTT 484
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              17 AGCCACGTCTCTACCTT 1
                                                                                                                                                                                                                                                                                                                                                                                                               19 eccaedardadedried 3
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Query Match
Best Local Similarity 94.1
Matches 16; Conservative
                                                                                                                                                                    воигсе
                                                                              검검점앙문료단
          PF
PR
PI
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  DEFINITION
ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 117
CQ867622
                                                                                                                                                                                                                         source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 116
CQ829824/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   FEATURES
                                                                                                                                                                                                       FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    VERSION
```

; 0

Gaps

.; 0

PAT 25-SEP-2002

8 셤

. 0

Gaps

.. 0

```
ö
                                                                                                                   PAT 21-JAN-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                PAT 29-SEP-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PAT 14-FEB-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                               Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                linear
                                                                                                                     linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1 (bases I to 20)
Benseler.F., Colley,J.L., Olsen,D.B. and Kuo,L.C.
Capped synthetic RNA, analogs, and aptamers
Patent: US 5861501-A 2 19-JAN-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match 0.4%; Score 15.2; DB 1; Best Local Similarity 85.0%; Pred. No. 1.1e+02; Matches 17; Conservative 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match

0.4%; Score 15.2; DB 1;
Best Local Similarity 85.0%; Pred. No. 1.1e+02;
Matchés 17; Conservative 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                DNA
                                                                                                                                                                                                                                                           Wagener, C.
METHOD FOR DETECTING MUTATED ALLELS
PACENT: WO 9839472-A 4 11-SEP-1998;
WAGENER CHRISTOPH (DE)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       20 bp L
Sequence 31 from patent US 6080546.
AR100074
                                                                                                                                                                                                                                                                                                                                                               /organism="unidentified"
/mol_type="unassigned DNA"
/db_xref="taxon:32644"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1. .20
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 2 from patent US 5861501.
AR030970
                                                                                                                20 bp
Sequence 4 from Patent WO9839472.
A86518
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  3173 TITIAAGICIGICICCTIAC 3192
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              2683 GAAGAGGAAAATAAAACC 2702
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1 rrrcaacrcrercrercrec 20
     1089 AATGITTCTTCATTTTC 1105
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20 GAAAATTAAAAATAAAACC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AR100074.1 GI:12810522
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AR030970.1 GI:5944184
                                                                                                                                                                 A86518.1 GI:6735117
                                 4 AATGATTCTTCATTTTC
                                                                                                                                                                                                                                                 (bases 1 to 20)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Unknown.
Unclassified.
                                                                                                                                                                                                                unidentified unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Unclassified.
                                                                                                                                                                                               unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                   source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DEFINITION
ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 123
AR030970/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AR100074/c
LOCUS
                                                                                   RESULT 122
A86518
                                                                                                                                DEFINITION
                                                                                                                                                                                                                ORGANISM
                                                                                                                                                ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 124
                                                                                                                                                                                                                                                                   AUTHORS
                                                                                                                                                                                                                                                                                                JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               JOURNAL
                                                                                                                                                                                                                                                 REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               KEYWORDS
                                                                                                                                                                                                                                                                                                                                FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                FEATURES
                                                                                                                                                                                                 SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              LOCUS
      ઢ
                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                  PAT 17-AUG-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            C12Q1/04, (C12Q1/68, C12R1:63), (C12Q1/04, C12R1:63), C12N15/00 CC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              OS Artificial Sequence
PN JP 2001340087-A/6
PD 11-DEC-2001
PF 31-MAY-2000 JP 2000166504
PI TETSUYA ISHIZUKA, TAKAHIKO ISHIGURO, JUICHI SAITO PC
GOLN13/58//
PC C1201/04, (C1201/68, G01N33/59, PC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       20 bp DNA linear PAT 27-AUG-2
Method for detecting thermostable hemolysin-analogous hemolysin
gene of Vibrio parahaemolyticus.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 other sequences, artificial sequences.

1 (bases 1 to 20)

Ishizuka,T., Ishiguro,T. and Saito,J.

Method for detecting thermostable hemolysin-analogous hemolysin gene of Vibrio parahaemolyticus
Patent: JP 2001340087-A 6 11-DEC-2001;
                                                                                                                                                                                              Unknown.
Unclassified.
1 (bases 1 to 20)
1 shizuka,T., Ishiguro,T. and Saitoh,J.
Oligomucleotides for detection of Vibrio parahaemolyticus and detection method for Vibrio parahaemolyticus using the same oligomucleotides
Patent: US 6562955-A 30 13-MAY-2003;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                             Query Match 0.4%; Score 15.4; DB 1; Length 20; Best Local Similarity 94.1%; Pred. No. 1e+02; Matches 16; Conservative 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match 0.4%; Score 15.4; DB 1; Length 20; Best Local Similarity 94.1%; Pred. No. 1e+02; Matches 16; Conservative 0; Mismatches 1; Indels

    .20
    /organism='Artificial Sequence'

                                                                                                    linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Location/Qualifiers

    .20
    /organism="synthetic construct"
|mol_type="genomic DNA"
|db_xref="taxon:32630"

                                                                                                    DNA
                                                                                             20 bp I
Seguence 30 from patent US 6562955.
AR317378
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1089 AATGTTTCTTCATTTTC 1105
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            BD084033.1 GI:22629643
JP 2001340087-A/6.
                                                                                                                                              AR317378.1 GI:33698472
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                4 AATGATICTICATITIC 20
19 TCATTTTCCTGGTGGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            synthetic construct
synthetic construct
                                                                                                                                                                              Unknown.
                                                                                                          DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            SOURCE
ORGANISM
                                                                                                                                                                                              ORGANISM
                                                              RESULT 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                               AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               RESULT 121
BD084033
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AUTHORS
TITLE
                                                                                                                                                                                                                              REFERENCE
                                                                                                                                                                                                                                                                                                              JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            JOURNAL
                                                                                 AR317378
                                                                                                                                                                                                                                                                                                                               FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            COMMENT
```

쉽

Page

FEATURES

à 엄

```
synthetic construct
constr
                                                                                                  BD184152 20 bp DNA linear PAT 17-JUN-2003
Method and detector for identifying subtypes of human papiloma
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         BD184313.1 GI:31876513

JP 2002360271-A/292.

Synthetic construct

SM synthetic construct

other sequences; artificial sequences.

I (bases 1 to 20)

E 1 (bases 1 to 20)

Ling, C., Lin, R., Yoo, Z., Huang, X., Lee, B., Lee, S., Lin, Y.,

Huang, C., Lin, Rau, H., Shi, C., Yeh, C., Cao, Y. and Pan, C.

Method and detector for identifying subtypes of human papiloma

L patent: JP 2002360271-A 292 17-DEC-2002;

KING CAR PROOD INDUSTRIAL CO LTD

OS Artificial Sequence

PD 17-DEC-2002

PD 17-DEC-2001

PP 28-NOV-2001 JP 2001362595
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            BD184313 20 bp DNA linear PAT 17-JUN
Method and detector for identifying subtypes of human papiloma
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 1.1e+02;
ttive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ce 1..20
/organism='Artificial Sequence'
Location/Qualifiers

    .20
        //organism="synthetic construct"
//organism="genomic DNA"
//db_type="genomic DNA"
//db_xref="taxon:32630"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           3304 ATTTTTTTTTTTTATATCC 3323
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20 Arrircarrirrarardrac 1
                                                                                                                                                                                   BD184152
BD184152.1 GI:31876352
JP 2002360271-A/131.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
Best Local Similarity
Matches 17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     viruses.
BD184313
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      RESULT 128
BD184313/c
LOCUS
                                                      RESULT 127
BD184152/c
                                                                                                       LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ORGANISM
                                                                                                                                                                                 ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                           ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           JOURNAL
                                                                                                                                                                                                                                                                                                                                              REFERENCE
                                                                                                                                                                                                                                                                                                                                                                    AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            COMMENT
                                                                                                                                                                                                                                                                                                                                                                                                                        TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    COMMENT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         PAT 20-APR-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  PAT 14-FEB-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1 (bases 1 to 20)
Bennett, C.Frank. and Vickers, T.A.
Bennett, C.Frank. and Vickers, T.A.
Oligonucleotide compositions and methods for the modulation of the expression of B7 protein
expression of B7 protein
Patent: US 6319906-A 128 20-NOV-2001;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   .
0
                                                                                                                                                                                                                                                                                             ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 1.1e+02;
iive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                       0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 1.1e+02;
tive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ch 0.4%; Score 15.2; DB 1; Length 20; 1 Similarity 85.0%; Pred. No. 1.1e+02; 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1 (bases 1 to 20)
Benseler,F., Cole,J.L., Olsen,D.B. and Kuo,L.C.
Capped synthetic RNA, analogs, and aptamers
Patent: US 6111095-A 2 29-AUG-2000;
Location/Qualifiers
  1 (bases 1 to 20)
Monia, B.P., Gaarde, W. and Cowsert, L.M.
Antisense modulation of MEKKS expression
Patent: US 6080546-A 31 27-UNN-2000;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20 bp DN Sequence 128 from patent US 6319906. AR178882 AR178882.1 GI:20220020
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1. .20
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1. .20 /organism="unknown" /mol_type="unassigned DNA"
                                                                                                                                                                                 /mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AR108815
Sequence 2 from patent US 6111095.
AR108815
AR108815.1 GI:12824302
                                                                                                       Location/Qualifiers
                                                                                                                                        1. .20
/organism="unknown"
                                                                                                                                                                                                                                                                                                                                              2236 AGCTAGTAAAGAATTTAGAA 2255
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     2683 GAAGAGAAAATAAAACC 2702
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1317 TTGAGTTTCAAAGGTTGCTG 1336
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1 rrradrircacadcirecte 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              GAAAATTAAAAATAAAAACC 1
                                                                                                                                                                                                                                                                                                                                                                            AGCTGGTAGAGACTTTAGAA 1
                                                                                                                                                                                                                                                                                           17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Unclassified.
                                                                                                                                                                                                                                                               Best Local Similarity
Matches 17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
Best Local Similarity
Matches 17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Unknown
                                                                                                                                                                                                                                                                                                                                                                                               20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
REFERENCE
AUTHORS
TITLE
JOURNAL
                                                                                                                                          source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   LOCUS
DEFINITION
ACCESSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Bource
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 125
AR108815/c
```

AUTHORS TITLE JOURNAL

FEATURES

REFERENCE

RESULT 126 AR178882

à g VERSION

REFERENCE AUTHORS TITLE

JOURNAL

FEATURES

ઠે

PAT 17-JUN-2003

ö

Gaps

PAT 07-0CT-1996

31

ö

Gaps

ö

```
Unclassified.
I (bases 1 to 20)
Hirschberg/C.B., Orellana,A., Hashimoto,Y., Swiedler,S.J., Wei,Z. and Ishihara,M.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Length 20;
                                                                                                                                                                                linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      linear
                            Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        linear
                                                                                                                                                                                                                                                                                                                   1 (bases 1 to 20)
Leonard, W.J., Noguchi, M. and McBride, O. Wesley.
Methods for diagnosis of XSCID and kits thereof
Patent: US 5518880-A 42 21-WAY-1996;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Glycosaminoglycan specific sulfotransferases
Patent: US 5541095-A 17 30-JUL-1996;
Location/Qualifiers
          85.0%; Pred. No. 1.1e+02; ive 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.4%; Score 15.2; DB 1;
85.0%; Pred. No. 1.1e+02;
iive 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0.4%; Score 15.2; DB 1;
85.0%; Pred. No. 1.1e+02;
tive 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DNA
                                                                                                                                                                                  DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      123988 20 bp 1
Sequence 17 from patent US 5541095.

    .20
    /organism="unknown"
    /mol_type="unassigned DNA"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                  121071 20 bp
Sequence 42 from patent US 5518880.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20 bp
Sequence 2 from patent US 6369208.
AR205764
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                2093 AGTATCTGTTGTAGCAGTTC 2112
                                                            3762 CAGATGGCTGGGATCCCTCC 3781
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1840 CCAAAAACAGGAACTACAG 1859
                                                                                           1 CACGIGGCIGGGAICGCICC 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1 AGAATCTGTTGTTCCAGTTC 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20 CCAGAAACAGGCACCACAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AR205764.1 GI:21503429
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      I23988.1 GI:1603858
                                                                                                                                                                                                                                    I21071.1 GI:1601425
       Best Local Similarity 85.0 Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match 0.4
Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                        Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Unknown.
                                                                                                                                                                                                                                                                    Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Unknown
                                                                                                                                                                                                                                                                                          Unknown
                                                                                                                                                                                                                    21071
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                              source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  source
                                                                                                                                                RESULT 130
121071
                                                                                                                                                                                                  DEFINITION
                                                                                                                                                                                                                                                                                    ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AR205764/c
                                                                                                                                                                                                                ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                      AUTHORS
TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 132
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 131
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                        REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      23988/c
                                                                                                                                                                                                                                                                                                                                                                                          FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                FEATURES
                                                                                                                                                                                                                                                                      SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          LOCUS
                                                                                                                                                                                  LOCUS
                                                                ઠે
                                                                                            셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Š
PR 04-rd.
PI CHING-YEE LING, ALL.
PI CHING-YEE LING, ALL.
PI HARG-LES, YI-JU LIN, CI-CHUNG HULL.
PI SHENG-LEE, YI-JU LIN, CI-CHUNG HULL.
PI SHENG-LEE, YI-JU LIN, CI-CHUNG HULL.
PI CHIH-XIN YEH, YI-FENG CAO, CHIH-LONG PAN
PC CI2N15/09, CI2N15/09, CI2N15/09, CI2N15/09, CI2N15/09, CI2N15/09, CI2N15/09, CI2N15/09, CI2N13/53, GOIN33/53, GOIN33/53, GOIN33/53, GOIN33/53, GOIN33/53, CI2Q1/70, CI2Q1/70, CI2Q1/70, CI2Q1/70, CI2Q1/70, CI2Q1/70, CI2N15/00, CI2N15/00
PC GOIN33/53, GOIN33/506, For identifying HPV 52. FH Key
Location/Qualifiers
//organism='Artificial Sequence'.
                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             E13800 20 bp DNA linear PAT 27-APR-1998 PCR primer for discriminating genotype 2a of HCV (Hepatitis C
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               C12N15/09, C07H21/04, C12Q1/68, C12Q1/70, (C12N15/09, C12R1:92); CC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              352511 PI
MIZOGAMI
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1 (bases I to 20)
CDO, T., Mukaide, M., Hikichi, K. and Mizogami, M.
NEW OLIGONUCLEOTIDE, PRIMER FOR DISCRIMINATION IN GENOTYPE OF
HEPATITIS C VIRUS COMPRISING THE SAME AND DISCRIMINATION IN
GENOTYPE OF HEPATITIS C VIRUS BY USING THE PRIMER
PATENT: JP 1997234072-A 52 09-SEP-1997;
                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    'organism='Artificial sequences' FT
                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       01-FEB-1996 JP 1996038875
01-FEB-1995 JP 95P 35997, 30-DEC-1995 JP 95P
TOMOYOSHI, MUKAIDE MASAKAZU, HIKICHI KAZUWASA, PI
                                                                                                                                                                                                                                                                                                                                                        Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%; Score 15.2; DB 1; Length 20;
                                                                                                                                                                                                                                                                                                                                                                                        3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     'note='Primer,OMM2aA'
                                                                                                                                                                                                                                                                                                                                                     0.4%; Score 15.2; DB 1;
85.0%; Pred. No. 1.1e+02;
tive 0; Mismatches 3;

    .20
    forganism="synthetic construct"
/mol type="genomic DNA"
    /db_xref="taxon:32630"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Location/Qualifiers

    .20
    /organism="unidentified"
    /mol_type="genomic DNA"
    /db_xref="taxon:32644"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                          3304 ATTITIATTITATATCC 3323
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Artificial sequences.
JP 1997234072-A/52
09-SEP-1997
                                                                                                                                                                                                                                                                                                                                                                                                                                                         20 ATTTTCATTTTTATGTGC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                strandedness: Single,
topology: Linear;
hypothetical: No;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                E13800
E13800.1 GI:3252568
JP 1997234072-A/52.
unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  anti-sense: No;
                                                                                                                                                                                                                                                                                                                                               Query Match
Best Local Similarity 85.09
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      misc_feature
FT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    unidentified
unclassified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              MASAFUMI
PC C12N
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                80
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                8888
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match
                                                                                                                                                                                                                                                                      source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 129
E13800
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        JOURNAL
                                                                                                                                                                                                                                                  FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          COMMENT
                                                                                                                                                                                                                                                                                                                                                                                                                          ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                         셤
```

ö

Gaps

ö

PAT 07-0CT-1996

PAT 20-JUN-2002

ö

Gaps

```
PAT 21-NOV-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             PAT 17-AUG-2003
                                                                                        PAT 12-JUN-2003
                                                                                                                                                                                                                   In (Dases I to 20)
Griffais,R., Hoiseth,S.K., Zagursky,R.J., Metcalf,B.J., Peek,J.A., Sankaran,B. and Fletcher,L.D. Chlamydia pneumoniae polymucleotides and uses thereof Patent: US 6559284-A 2018 06-MAY-2003;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Antisense modulation of Interferon gamma receptor 1 expression patent: US 6566132-A 67 20-MAY-2003;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Barany, F., Zirvi, M., Gerry, N.P., Favis, R. and Kliman, R.
                                                                                                                                                                                                                                                                                                                                                                                                        0
                                                                                                                                                                                                                                                                                                                                                                       Query Match

0.4%; Score 15.2; DB 1; Length 20;
Best Local Similarity 85.0%; Pred. No. 1.1e+02;
Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match
0.4%; Score 15.2; DB 1; Length 20;
Best Local Similarity 85.0%; Pred. No. 1.1e+02;
Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20 bp Dh Sequence 6843 from Patent WO0179548. AX295081.1 GI:17056764
                                                                                        AR311481 20 bp DN Sequence 2018 from patent US 6559294.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AR337006 20 bp
Sequence 67 from patent US 6566132.
AR337006
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1. .20
/organism="unknown"
/mol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                                 1. .20
/organism="unknown"
/mol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1689 GAGCAGTTGTCTCCAACAGC 1708
                                                                                                                                                                                                                                                                                                                                                                                                                                       1617 GAAAACTTCCTTCCAGATAT 1636
                                                                                                                                                                                                                                                                                                                                                                                                                                                         20 GAAAACTTCCCTCCAGTCAT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    20 GAGCCGTTGTCTCCAGCAAC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AR337006.1 GI:33722860
                                                                                                                             AR311481
AR311481.1 GI:31704907
20 TTGACAGAAACCACATTCA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (bases 1 to 20)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Unclassified.
                                                                                                                                                                                          Unknown.
Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Watt, A.T.
                                                                                                                                                                     .
Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 137
AX295081
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AR337006/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            REFERENCE
AUTHORS
TITLE
JOURNAL
FEATURES
                                                               RESULT 135
AR311481/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DEFINITION
                                                                                                             DEFINITION
                                                                                                                                                                                        ORGANISM
                                                                                                                                                                                                                                                                    TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RESULT 136
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
                                                                                                                            ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                       REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      d
                                                                                                                                                                                                                                                                                                                                                                                                                                              ઠે
                     셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          PAT 12-JUN-2003
                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                    PAT 20-DEC-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gatti.R.A.
Modified SSCP method using sequential electrophoresis of multiple nucleic acid segments
Patent: US 6458536-A 93 01-OCT-2002;
Location/Qualifiers
                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                            .
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 1.1e+02; Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Length 20;
                                                                                                                                                                          Query Match
0.4%; Score 15.2; DB 1; Length 20;
Best Local Similarity 85.0%; Pred. No. 1.1e+02;
Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                     linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Indels
 M Unknown.
Unclassified.
3 1 (bases 1 to 20)
S Cole, Ji., Kuo, L.C., Olsen, D.B. and Benseler, F.
Capped synthetic RNA, analogs, and aptamers
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match

0.4%; Score 15.2; DB 1;
Best Local Similarity 85.0%; Pred. No. 1.1e+02;
Matches 17; Conservative 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                     DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Antisense modulation of raidd express:
Patent: US 6551826-A 151 22-APR-2003;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AR307940 20 bp D1
Sequence 151 from patent US 6551826.
AR307940.1 GI:31698696
                                                                                                                    1. .20
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                  AR233731 20 bp 1
Sequence 93 from patent US 6458536.
AR233731 GI:27276355
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1. .20
/organism="unknown"
/mol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       /organism="unknown"
/wol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1989 TGTGCAACACCTTCAGATAA 2008
                                                                                                                                                                                                                                         2683 GAAGAGGAAAATAAAACC 2702
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         489 TTGACAGGAAACCCCATCCA 508
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1 TTTGCAACACCTTCACCTAA 20
                                                                                                                                                                                                                                                                         20 GAAATTAAAAATAAAACC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Unknown.
Unclassified.
1 (bases 1 to 20)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (bases 1 to 20)
                                                                                                                                                                                                                                                                                                                                                                                                                                                 Unknown.
Unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Watt, A.T.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                               Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 134
AR307940/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     source
                                                                                                                                                                                                                                                                                                                                                                    DEFINITION
ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DEFINITION
ACCESSION
                                                                                                                     source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AUTHORS
TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                 ORGANISM
           ORGANISM
                                                                    TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                                        RESULT 133
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                  KEYWORDS
                                        REFERENCE
                                                     AUTHORS
                                                                                                                                                                                                                                                                                                                                          AR233731
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             FEATURES
```

셤 ò

ó

Gaps

; 0

Gaps

10001863-3.81.rge

```
/mol_type="genomic DNA"
/db_xref="taxon:32630"
/note="Oligonucleotide for Identifying HPV 33"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20 ATTTCATTTTATATGTGC 1
                                                                                                                                                                  20 Arrrrcarrrrarargrac 1
                                                                                                                                                                                                                                                                                                 AX742460.1 GI:30576428
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AX962807.1 GI:40881920
                                                                                                                                                                                                                                                                                                                                      synthetic construct
synthetic construct
                                                           Ouery Match
Best Local Similarity 85.09
Matches 17, Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AX962807
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AX962807
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            viruses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         source
                                                                                                                                                                                                                   RESULT 140
AX742460/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 141
AX962807/c
                                                                                                                                                                                                                                                                                                                                      SOURCE
ORGANISM
                                                                                                                                                                                                                                                                  DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                     ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                         REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                        AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    JOURNAL
                                                                                                                                                                                                                                                                                                                     KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      PEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                         TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           a
                                                                                                                                  ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              8
                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
Method of designing addressable array for detection of nucleic acid
                                                                                                                                                                                                                                                                                                                                                                       PAT 23-NOV-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   PAT 12-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Lin,C.Y., Lin,R.W., You,C.M., Huang,H.H., Lee,B.H., Lee,H.H., Lin,Y.'J., Pan,C.C., Hsu,H.C., Shih,C.W., Yeh,C.H., Kao,Y.F., Pan,C.L. and Chan,P.
Method and detector for identifying subtypes of human papilloma
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Spagnoli, R., Achstetter, T., Cauet, G., Degryse, E., Dumas, B., Pompon, D. and Winter, J.
Fast strains autonomously producing steroids
Patent: WO 02061109-A 45 08-AUG-2002;
Aventis Pharma S.A. (FR)
                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
               sequence differences using ligase detection reaction
Patent: WO 0179548-A 6843 25-OCT-2001;
CORNELL RESEARCH FOUNDATION, INC. (US)
                                                                                                                                                                                Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 1.1e+02; Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 1.18+02;
ive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     linear
                                                                                                                                                                                                                                                                                                                                                                         linear
                                                                              1.20
/organism="synthetic construct"
/mol_type="unassigned DNA"
/db_xref="taxon:32630"
/note="Hypothetical Probe Sequence"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1. .20
/organism="synthetic construct"
/mol_type="unassigned DNA"
/db_xref="taxon:32630"
/note="Oligonucleotide leu2D"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       organism="synthetic construct"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Patent: EP 1302550-A 131 16-APR-2003;
King Car Pood Industrial Co., Ltd. (TW)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                20 bp I
Sequence 45 from Patent WO02061109.
AX544171
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            20 bp | Sequence 131 from Patent EP1302550.
                                                                Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Location/Qualifiers
                                                                                                                                                                                                                                                    1346 TGATTTTGGGACAACCAGCC 1365
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1558 GCATCTTCAATGGCTTGTCC 1577
                                                                                                                                                                                                                                                                     1 TGATTCTGGGACCAGCAGCC 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20 GCATCTTCAATGGCCTTACC 1
                                                                                                                                                                                                                                                                                                                                                                                                                      AX544171.1 GI:25277733
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AX742328.1 GI:30576296
                                                                                                                                                                                                                                                                                                                                                                                                                                                        synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
Best Local Similarity
Matches 17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                  DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                  Bource
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Bource
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       source
                                                                                                                                                                                                                                                                                                                                                    AX544171/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AX742328/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ORGANISM
                                                                                                                                                                                                                                                                                                                                      RESULT 138
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TITLE
JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        REPERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      JOURNAL
                               JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AUTHORS
                                                                  FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       PEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       TITLE
TITLE
                                                                                                                                                                                                                                                    ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        g
```

```
ö
                                                                                                                                                                                                                                                                          PAT 12-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            PAT 14-JAN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Lin,C.Y., Lin,R.W., You,C.M., Huang,H.H., Lee,B.H., Lee,H.H., Lin,Y.-J., Fan,C.C., Hsu,H.C., Shih,C.W., Yeh,C.H., Kao,Y.F., Pan,C.L. and Chan,P.
Method and detector for identifying subtypes of human papilloma
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Baker, B.F., Freier, S.M. and Dobie, K.W.
Antisense modulation of il-1 receptor-associated kinase-1
expression
Patent: Word 130458-A 63 18-DEC-2003;
ISIS PHARMACEUTICALS, INC. (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /mol_type="genomic DNA"
/db_xref="taxon:32630"
/note="Oligonucleotide for Identifying HPV 52"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 1.1e+02; Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1. .20
// Organism="synthetic construct"
// Mol_type="unassigned DNA"
// Ab_xref="taxon:32630"
// noTe="Antisense Oligonucleotide"
0.4%; Score 15.2; DB 1;
85.0%; Pred. No. 1.1e+02;
tive 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match

0.4%; Score 15.2; DB 1;
Best Local Similarity 85.0%; Pred. No. 1.1e+02;
Matches 17; Conservative 0; Mismatches 3;

    .20
    /organism="synthetic construct"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Patent: EP 1302550-A 263 16-APR-2003;
King Car Food Industrial Co., Ltd. (TW)
Location/Qualifiers
                                                                                                                                                                                                                                                                          DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                 other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  synthetic construct
synthetic construct
other sequences; artificial sequences.
                                                                                                                                                                                                                                                                20 bp 1
Sequence 263 from Patent EP1302550.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AX962807 20 bp
Sequence 63 from Patent WO03104458.
                                                                                                 3304 ATTITITITITIATATCC 3323
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   3304 ATTITITITITITITITIC 3323
```

qq ઠે

10001863-3.sl.rge

```
Regulation of repressor genes using nucleic acid molecules. BD256407 BD256407. GI:33066177 JP 2002541795-A/4200. unidentified unidentified unclassified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  PAT 29-SEP-1999
                                                                                                        PAT 05-DEC-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gaps
                                                                                                                                                                                                 Unknown.
Unclassified.
1 (bases I to 17)
3 (bases I to 17)
Glucksmann,M.Alexandra.
Methods for identifying compounds useful in treating type II diabetes
Alent: US 5795726-A 24 18-AUG-1998;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Unknown.
Unclassified.
Unclassified.
Glucksmann, M. Alexandra.
Assays for diagnosing type II diabetes in a subject
Patent: US 5800998-A 21 01-SEP-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.4%; Score 15; DB 1; Length 17;
100.0%; Pred. No. 89;
tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                         Query Match 0.4%; Score 15; DB 1; Length 17; Best Local Similarity 100.0%; Pred. No. 89; Matches 15; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    linear
                                                                                                         linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               17 bp I
Sequence 21 from patent US 5800998.
AR036967
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1. .17
/organism="unknown"
/mol_type="unassigned DNA"
                                                                                                        AR023742 17 bp 1
Sequence 24 from patent US 5795726.
AR023742 AR023742.1 GI:3977036
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Location/Qualifiers
AR036967.1 GI:5954823
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      402 CTGCAGGTGCTGGAT 416
                                                                                                                                                                                                                                                                                                                                                                                                                                                           402 CTGCAGGTGCTGGAT 416
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         2 CTGCAGGTGCTGGAT 16
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               2 CTGCAGGTGCTGGAT 16
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match 0.4
Best Local Similarity 100.
Matches 15; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Unknown.
                                                                                                                                                                                         Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AUTHORS
TITLE
JOURNAL
FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                BD256407/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DEFINITION
ACCESSION
                                                                                                              LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                      source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 145
AR036967
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 146
                                                                                 RESULT 144
AR023742
                                                                                                                                                                                                           ORGANISM
                                                                                                                                                                                                                                                        AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                      JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 REFERENCE
                                                                                                                                                                                                                                         REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           δ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         PAT 27-AUG-2002
                                                                                                                  PAT 14-JAN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Description of the Artificial sequence: oligonucleotides FH Location/Qualifiers source 1. .20
                                                                                                                                                                                                                        Eukaryota, Metazoa, Chordata, Craniata, Vertebrata, Euteleostomi,
Mammalia, Eutheria, Primates, Catarrhini, Hominidae, Homo.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                        Baker, B.F., Freier, S.M. and Dobie, K.W.
Antisense modulation of il-1 receptor-associated kinase-1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ó
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match

0.4%; Score 15.2; DB 1; Length 20;
Best Local Similarity 85.0%; Pred. No. 1.1e+02;
Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    /organism='Artificial Sequence'
                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 1.1e+02; Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1. .20
Corganism="synthetic construct"
/mol_type="genomic DNA"
/db_xref="taxon:32630"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Wagner, C.
Method for detecting mutated alleles
Patent: JP 2001514504-A 4 11-SEP-2001;
CHRISTOPH WAGNER
OS Artificial Sequence
PN JP 2001514504-A/4
PD 11-SEP-2001
PF 04-MAR-1999 DF 1998538071
PR 04-MAR-1997 DE 197 08 758.2
PI CHRISTOPH WAGNER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20 bp DNA Method for detecting mutated alleles. BD070557
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DD070557.1 GI:22616160
JP 2001514504-A/4.
synthetic construct
synthetic construct
other sequences; artificial sequences.
1 (bases 1 to 20)
                                                                                                                                                                                                                                                                                                        expression
Patent: WO 03104458-A 129 18-DEC-2003;
ISIS PHARMACEUTICALS, INC. (US)
Location/Qualifiers
                                                                                                                       DNA
                                                                                                                     AX962873 20 bp DN
Sequence 129 from Patent WO03104458.
AX962873.1 GI:40881996

    .20
    /organism="Homo sapiens"
    /mol_type="unassigned DNA"
    /db_xref="taxon:9606"

                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    460 AGAGCCTAAGCCACCTCTCT 479
           460 AGAGCCTAAGCCACCTCTCT 479
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1 AGAGCCTAGGCGGCCTCTCT 20
                               20 AGAGCCTAGGCGGCCTCTCT 1
                                                                                                                                                                                                 Homo sapiens (human)
                                                                                                                                                                                                                     Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               source
                                                                                                                                                   ACCESSION
VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                     AX962873
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 143
BD070557
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AUTHORS
TITLE
JOURNAL
                                                                                                                                                                                                                                                               REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        REFERENCE
                                                                                                                                                                                                                                                                                                                             JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               FEATURES
                                                                                                                                                                                                                                                                                                                                                             FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        COMMENT
```

;

35

ö

REFERENCE AUTHORS TITLE JOURNAL

COMMENT

```
PAT 07-SEP-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PAT 07-SEP-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Blatt, L., Mcswiggen, J. and Chowrira, B.M.
Method and reagent for the modulation and diagnosis of cd20 and nogo gene expression
Patent: WO 0159103-A 2853 16-AUG-2001;
RIBOXYME PHARMACEUTIAALS, INC. (US); Blatt, Lawrence (US);
McSwiggen, James (US); Chowrira, Bharat M. (US)
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Blatt, L., Mcswiggen, J. and Chowrira, B.M.
Method and reagent for the modulation and diagnosis of cd20 and nogo gene expression
Patent: WO 0159103-A 3219 16-AUG-2001;
RIBOXYME HARMACEUTICALS, INC. (US) ; Blatt, Lawrence (US) ;
McSwiggen, James (US) ; Chowrira, Bharat M. (US)
                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Score 15; DB 1; Length 17;
Pred. No. 89;
                                                                                                                                                                                             Length 17;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Length 17;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              linear
                                                                                                                                                                                                                               0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                  linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Indela
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1.17
/organism="synthetic construct"
/mol_type="unassigned RNA"
/db_xref="teaxon:32630"
/note="Nucleic Acid"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%; Score 15; DB 1;
ilarity 100.0%; Pred. No. 89;
Conservative 0; Mismatches (
                                                                                                                                                                                           Score 15; DB 1;
Pred. No. 89;
                                                                                                                                                                                                                                                                                                                                                                                                  RNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       synthetic construct
synthetic construct
other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                        17 bp RN Sequence 2853 from Patent WO0159103.
AX217411
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       AX217777
Sequence 3219 from Patent WOO159103.
AX217777
                                                                                                                                                                          Query Match 0.4%; Score 15; UB
Best Local Similarity 100.0%; Pred. No. 89;
Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match 0.4%; Score 15; DB Best Local Similarity 100.0%; Pred. No. 89; Matches 15; Conservative 0; Mismatches
                                                                                                                  /organism="unidentified"
                                                                                                                                   /mol_type="genomic DNA"
/db_xref="taxon:32644"
                                                                                                                                                                                                                                                                                                                                                                                                                                                     AX217411.1 GI:15527472
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AX21777.1 GI:15527838
                                                                                                                                                                                                                                                                   1896 CAGAATGACTTTGCT 1910
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                3797 CTGACAGGAGAACTA 3811
                                                                                                                                                                                                                                                                                      15 CAGAATGACTTTGCT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    16 crcacacacacaacra 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Local Similarity
nes 15; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
                                                                                                                                                                                                                                                                                                                                                                                                             DEFINITION
ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT 149
AX217777/c
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                             RESULT 148
AX217411/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Best Loca
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          KEYWORDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             FEATURES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                đ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            I7 bp DNA linear PAT 17-JUL-2003 BD256855 BD256855 I GI:33066625 UP 2002541795-A/4648.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Regulation of repressor genes using nucleic acid molecules FH Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Regulation of repressor genes using nucleic acid molecules FH Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      C12P21/02, C12P21/02//A61K31/711, (C12N5/10, C12R1:91), (C12P21/02, PC
                                                                                                                                                                                                                                                                                                                                                                            C12P21/02,C12P21/02//A61K31/711,(C12N5/10,C12R1:91),(C12P21/02, PC
                                                                                                                                                                                                                                                                                                                                                                                                           C12P21/02, C12R1:91), (C12P21/02, C12R1:91), C12N15/00, C12N5/00,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (C12P21/02,C12R1:91), (C12P21/02,C12R1:91),C12N15/00,C12N5/00,
A61K37/02,
(C12N5/00,C12R1:91)
                                                                                                                                                                                                                                                                                                        ы
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        11-APR-2000 JP 2000611654
12-APR-1999 US 60/129390
LAWRENCE BLATT,MICHAEL ZWICK,PAMELA PAVCO,JAMES MCSWIGGEN PC
5/09,A61K38/00,A61K48/00,A61P43/00,A61P43/00,C12N5/10, PC
                                                                                                                                                                                                          OS Eukaryote
PN JP 2002541795-A/4200
PD 10-DEC-20002
PP 11-ARR-2000 JP 2000611654
PR 12-APR-1999 US 60/129390
PI LAWRENCE BLATT,MICHAEL ZWICK,PAMELA PAVCO,JAMES MCSWIGGEN FC12N15/09,AG1R38/00,AG1R48/00,AG1R43/00,AG1R5/10,PC
THORS Blatt, L., Zwick, M., Pavco, P. and Mcswiggen, J.
TLE Regulation of repressor genes using nucleic acid molecules
TNAL Patent: JP 2002541795-A 4200 10-DEC-2002;
T OS Eukaryote
T NP JP 2002541795-A/4200
PD JO-DEC-2002
PF 11-APR-2007
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       unclassified.

1 (bases 1 to 17)
Blatt,L., Zwick,M., Pavco,P. and Mcswiggen,J.
Regulation of repressor genes using nucleic acid molecules Patent: JP 2002541795-A 4648 10-DEC-2002;
RIBOZYME PHARMACEUTICALS INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Length 17;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /organism='Eukaryote'.
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          /organism='Eukaryote'
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.4%; Score 15; DB 1;
100.0%; Pred. No. 89;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                100.0%; Pred. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1. .17
/organism="unidentified"
/mol_type="genomic DNA"
/db_xref="taxon:32644"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Eukaryote
JP 2002541795-A/4648
10-DEC-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                A61K37/02,
(C12N5/00,C12R1:91)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 1896 CAGAATGACTTTGCT 1910
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             15 CAGAATGACTTTGCT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Best Local Similarity 100.0
Matches 15; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             source
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        source
                                                                                                                                                                                                                                                                                                                                      C12P21/02,
PC
                                                                                                                                                                                                                                                                                                                                                                                                  C12R1:91)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
```

DEFINITION ACCESSION VERSION KEYWORDS SOURCE ORGANISM

REFERENCE AUTHORS TITLE

COMMENT

RESULT 147 BD256855/c

ઠે 셤 LOCUS

source

FEATURES

ö

ö

source

PEATURES

```
Unknown.
            Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                        Unknown.
                                                                                                                                                                                                                                                                                                                                                              DEFINITION
ACCESSION
VERSION
KEYWORDS
                                                                                                                                source
                                                                                                                                                                                                                                                                                                                   RESULT 153
AR226207
LOCUS
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                       SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             JOURNAL
FEATURES
                                                                                                   JOURNAL
FEATURES
                                                      REFERENCE
AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ò
                                                                                                                                                                                                                                                 ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           PAT 25-SEP-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  PAT 01-SEP-2000
                                                                                                                  PAT 07-SEP-2001
                                                                                                                                                                                                                                              Blatt, L., Mcswiggen, J. and Chowrira, B.M.
Method and reagent for the modulation and diagnosis of cd20 and nego gene expression
Patent: WO 0159103-4 3220 16-AUG-2001;
RIBOZYME PHARMACEUTICALS, INC. (US); Blatt, Lawrence (US);
McSwiggen, James (US); Chowrira, Bharat M. (US)
Location/Qualifiers
1. 17
/organism="synthetic construct"
/mol_type="unassigned RNA"
/db_xref="taxon:32630"
/note="Nucleic Acid"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Unclassified.

1 (bases 1 to 20)
Monia, B.P.
Antisense oligonuclectide modulation of B-raf gene expression
Patent: US 5981731-A 26 09-NOV-1999;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           linear
                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match 0.4%; Score 15; DB 1; Length 17; Best Local Similarity 100.0%; Pred. No. 89; Matches 15; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ch 0.4%; Score 15; DB 1; Length 20; ll Similarity 100.0%; Pred. No. 1.1e+02; 15; Conservative 0; Mismatches 0; Indels
                                                                                                                    linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DNA
                                                                                                                                                                                                                    other sequences; artificial sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AR216068 20 bp DN
Sequence 115 from patent US 6410518.
AR216068.1 GI:23314356
                                                                                                                  AX217778 17 bp Ri
Sequence 3220 from Patent WO0159103.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AR085548 20 bp 1
Sequence 26 from patent US 5981731.
AR085548.1 GI:10012315
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /mol_type="unassigned DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /organism="unknown"
                                                                                                                                                            AX217778.1 GI:15527839
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1467 AAACAAATGAGTGAG 1481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 3797 CTGACAGGAGACTA 3811
                  3797 CTGACAGGAGAACTA 3811
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      CTGACAGGAGAACTA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20 AAACAAATGAGTGAG 6
                                           17 CTGACAGGAGAACTA 3
                                                                                                                                                                                         synthetic construct
synthetic construct
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Best Local Similarity
Matches 15; Conserva
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Unknown.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Unknown
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              15
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 152
AR216068/c
LOCUS
DEFINITION
ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         KEYWORDS
SOURCE
ORGANISM
                                                                                                     AX217778/c
LOCUS
DEFINITION
ACCESSION
VERSION
KEYWORDS
SOURCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               DEFINITION
ACCESSION
VERSION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      RESULT 151
AR085548/c
LOCUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             source
                                                                                                                                                                                                        ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              TITLE
JOURNAL
                                                                                         RESULT 150
                                                                                                                                                                                                                                                  AUTHORS
TITLE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    REFERENCE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AUTHORS
                                                                                                                                                                                                                                     REFERENCE
                                                                                                                                                                                                                                                                                             JOURNAL
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            FEATURES
                                                                                                                                                                                                                                                                                                                                         FEATURES
                                           셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           g
                  ò
```

```
ö
                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                     PAT 20-DEC-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                       Gaps
Unknown.
Unknown.
Unclassified.
1 (bases 1 to 20)
Monial.B.P.
Antisense oligonucleotide inhibition of raf gene expression
Patent: US 6410518-A 115 25-JUN-2002;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ;
0
                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                         Query Match 0.4%; Score 15; DB 1; Length 20; Best Local Similarity 100.0%; Pred. No. 1.1e+02; Matches 15; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match 0.4%; Score 15; DB 1; Length 20; Best Local Similarity 100.0%; Pred. No. 1.1e+02; Matches 15; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Unclassified.

1 (bases 1 to 20)

Ward,D.T. and Watt,A.T.
Antisense modulation of helicase-moi expression
Patent: US 644466-A 88 03-SEP-2002;
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                         DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                     20 bp | 1
Sequence 88 from patent US 6444466.
AR226207
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Search completed: February 16, 2005, 09:21:47 Job time : 8 secs
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1. .20
/organism="unknown"
/wol_type="genomic DNA"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AR226207.1 GI:27264361
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              2331 ATCCATGAAGGTTTC 2345
                                                                                                                                                                                                                                                                                                               1467 AAACAAATGAGTGAG 1481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     4 Arccardadgerrrc 18
                                                                                                                                                                                                                                                                                                                                                   20 AAACAAATGAGTGAG 6
```

Human TLR4 gene ex Human TLR4 gene ex Human TLR4 gene am Human TLR4 gene ex	TLR4 gene TLR4 gene	Human TLR4 gene ex	ene	Human TLR4 gene ex Human TLR4 gene ex Human TLR4 gene ex	ene	ene er d	TLR4 gene TLR4 gene	TLR4 gene TLR4 gene	Human TLR4 gene ex Human TLR4 gene ex Human TLR4 gene ex	TLR4 gene TLR4 gene	TLR4 gene TLR4 gene	4.4.	CD14 monocl	Toll-lik	Toll-like	Toll-like Toll-like	k k	Toll-like Toll-like	Toll-like Toll-like	y k	Toll-like	Toll-like	Toll-like	Toll-like Toll-like	Human Toll-like re Human Toll-like re	Toll-like r	Toll-like r	Toll-like Toll-like	Toll-like	Toll-like	TOIL-1 tis E	Human TLR4 gene ex Human toll-like re	oll-like	111	C. albicans specif Human TLR4 gene ex	ув с
20 0.5 20 1 AAC84798 20 0.5 20 1 AAC84821 20 0.5 20 1 AAC84775 20 0.5 20 1 AAC84775	0.5 20 1	0.5 20 1	0.5	0.5 20 1	0.5 20 1	0.5 20 1	0.5 20 1	0.5 20 1	0.5 20 1 0.5 20 1 0.5 20 1	0.5 20 1 0.5 20 1	0.5 20 1	0.5 20 1	0.5	0.5	0.5 20 1	0.5 20 1	0.5 20 1	0.5 20 1	0.5 20 1	0.5 20 1	0.5	0.5	0.5 20 1	0.5 20 1	0.5 20 1	0.5	0.5 20 1	0.5 20 1	0.5 20 1	0.5	20 0.5 20 I .2 0.5 24 I	0.5 19 1	19 0.5 19 1	.4 0.5 24 1	.2 0.5 24 1 18 0.5 18 1	4 0.5 22 1
0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	8 6 F		7 <b>7</b> 7		47		52			5 8 5 9		63		66	3 6 9		72 73	74 75	76 77	78	80	85	84	82 86	88	6 B	91	83	94 70	96	97 98 19	99	101	02 18 03 18	104 18 105	06 17
version 5.1.6 - 2005 Compugen Ltd.	model .	3:05; Search time 15 Seconds	(Without all million cell updates/sec		totcactgacaggagaacta 3811			chosen parameters: 704			ıries		by chance to hav	<ul> <li>to the score of the result being printed, of the total score distribution.</li> </ul>	SUMMARIES			Human leuko 399Reverse			Human TLR4 Human TCR4	Human TLR	Human TL	Anti-CD14 Sense PCR	Human Human	Human TL	Human TL	Antisense PCR pi Human TLR4 gene	Human TLI	Human filt VEGF	Human TLK4 gene Human TLR4 gene	Human TLR4	Human toll-like	Probe for detect Human Toll like	Human TLR Human TLR	Human TLR4 g
GenCore ver Copyright (c) 1993 - 2	OM nucleic - nucleic search, using sw mo	Run on: February 16, 2005, 09:23:0		itle: erfect score:	 0)	Scoring table: IDENTITY NUC Gapext 0.5.	Searched: 352 seqs, 7150 residues	Total number of hits satisfying chosen p	Minimum DB seq length: 8 Maximum DB seq length: 80	Post-processing: Minimum Match 0%	Maximum Match 100% Listing first 352 summaries	Database : rngdb:*	No. is the number of	score greater than or equal to the and is derived by analysis of the		Query	Score Match Length DB	1 50 1.3 50 2 30 0.8 30	0.8 30 1 0.8 31 1	5 28.4 0.7 30 1 6 27 0.7 27 1	0.7 26 1	9 24 0.6	11 24 0.6 24 1	24 0.6 24	24 0.6 24 1 23 0.6 23 1	23 0.6 23	23 0.6 23 1	19 22.4 0.6 24 20 22 0.6 22	22 0.6 22 1	21.2 0.6 27 1	24 21 0.6 21 1 25 21 0.6 21 1	26 21 0.6 21 1 27 21 0.6 21 1	21 0.6 25 1	30 20 0.5 20 1		20 0.5 20 1

Human glucocortico Human HMG-CoA redu Oligonucleotide as Oligonucleotide as Human B7.1 DNA ant Primer #2 for iden Human Pas antisens Human oligonucleot Human ADAM15 targe Human Oligonucleot Human ADAM15 artisens Human ADAM15 artisens Human ADAM15 artisens Human ADAM15 artisens Human TWF alpha antisens Human TWF alpha antisense oligonucleot Human TWF alpha antisense oligonuc Oligo specific for Muuse MSP PCR primer, Oligo specific for Muuse MSP PCR primer, Pollen allergen ge Human COL9A2 PCR p NOV72 PCR primer, Pollen allergen ge Human C-myb hammer Single nucleotide Single nucleotide Single nucleotide WNV Inozyme substr HBV Inozyme substr HBPAILILIS B VITUS HEPAILILIS B VITUS HEPAILIS B VITUS HUMAN THR-AIDHA HA	
	4 18 1 AANS11819 4 19 1 AANS11819 4 20 1 AAANS1098 7 AANS1098 7 AANS11819 7 AANS1181
**************************************	C C C C C C C C C C C C C C C C C C C
tric tric tric tric tric borrange borran	Mouse casean kinas Human biallalic po Human biallalic po Human Toll-like re Endogenous caroten Antisense siNA that do Human Toll-like siNA oli Human TNF siNA oli Human apolipoprote Modifide oligonucl DCOH (PCBD) gene e Human apolipoprote Human apolipoprote Modifide oligonucl DCOH (PCBD) gene e Human B7-1 targett Human B7-1 targett Human B7-1 targett Human mdm2 phospho PCR primer for mar Human mdm2 phospho Human mdm2 antisen Human mdm2 antisen Human ndm2 antisen Human oligonucleot Human staniocaliy re M. gallisepticum m plant retroelement Human staniocalci Human staniocalci Human staniocalci Human glucocortico Human glucocortico
	20 1 ABS65155 19 1 AAXO9330 19 1 AC707938 19 1 ADF78593 19 1 ADF785738 19 1 ADF785714 19 1 ADG34714 19 1 ADG34714 19 1 ADG34714 19 1 ADG34714 19 1 ADG34714 19 1 ADG34714 19 1 ADG34714 20 1 AAX58720 20 1 AAX58720 20 1 AAX58720 20 1 AAX58720 20 1 AAX58720 20 1 AAX58720 20 1 AAX8672 20 1 AAX8672 20 1 AAX8672 20 1 AAX8672 20 1 AAX8672 20 1 AAX8672 20 1 ABZ89870 20 1 ABZ89
	4.4.4.4.4.4.4.4.4.4.4.4.4.4.4.4.4.4.4.

326 15.2 0.4 20 1 ADP49620 327 15.2 0.4 20 1 ADP49620 328 15.2 0.4 20 1 ADP84310 330 15.2 0.4 20 1 ADS2657 331 15.2 0.4 20 1 ADS2657 332 15.2 0.4 20 1 ADS0671 333 15.2 0.4 20 1 ADT00664 334 15.0 0.4 17 1 AAV41419 356 15 0.4 17 1 AAV41419 36 15 0.4 17 1 AAV6094 37 15 0.4 17 1 AAV6097		RESULT 1 ABZ02491 ID ABZ02491 standard; DNA; 50 BP. XX AC ABZ02491; XX DT 09-JAN-2003 (first entry) XX XX DT 09-JAN-2003 (first entry) XX	T7; leukocyte; gne expression profiling; allograft reject atherosclerosis; congestive heart failure; systemic lupus rheumatoid arthritis; osteoarthritis; cytomegalovirus; inf 88.  Homo sapiens.  WO200257414-A2. 25-JUL-2002.	ед е е со о и з
Human ATM gene exc Human nucleolin ph Mouse RAIDD antise Human lipoprotein Autonomously stero Capture oligonucle Human CREB phospho Vitamin D nuclear Antinsense oligonu Human papillomavir Human papillomavir Human papillomavir	Auman B7-1 targete Human B7-1 targete HPV 33 detecting p HPV 52 detecting p Single nucleotide Human oligonucleot Human oligonucleot Human oligonucleot Human oligonucleot Human noligonucleot Human noligonucleot	Human cathepein C- Human MCP4 derived Human ICAM-derived AA284245-derived AA497002-derived o AA497002-derived o AA497002-derived o Human SREBP-1 anti Human SREBP-1 targ Human glucocortico Human glucocortico Human IRAK-1 DNA Human IRAK-1 DNA	Cyclin dependent k Cyclin dependent k Human polo-like ki Human polo-like ki Human requiem anti Oligonucleotide as Oligonucleotide as Oligonucleotide as Human resistin ant Human B7-1 DNA ant Human B7-1 DNA ant Mouse complement c Human endothelial	enderic lic lic lic lic lic lic lic lic lic l
				20 1 ADJ21649 20 1 ADJ24576 20 1 ADK79428 20 1 ADK73906 20 1 ADK73906 20 1 ADK81604 20 1 ADK81604 20 1 ADK81604 20 1 ADK7411 20 1 ADK7411 20 1 ADK7411 20 1 ADK7411 20 1 ADK75910 20 1 ADC45910 20 1 ADP31773 20 1 ADP31773 20 1 ADP31773
553 554 555 556 557 557 559 60 60 60 60 60 60 60 60 60 60 60 60 60	2666 2666 2666 2666 271 271 272 273 274 275 276 277 277 277 277 277 277 277 277 277	888 888 888 888 888 888 888 888 888 88	9985 998 998 998 998 998 998 998 998 998	307 C 308 310 310 311 312 313 313 314 315 315 316 317 318 316 317 318 319 319 319 319 319 319 319 319

```
gene, where the gene corresponds to any of 8143 oligonucleotides (ABZ00010-ABZ08152) each having 50 base pairs (bp). The system is useful dispussion syrpression profiling It is particularly useful for dispussing a disease, monitoring (rate of) progression of a disease, predicting therapeutic outcome, determining prognosis for a patient, predicting disease complications in an individual or monitoring response to treatment in an individual. The diseases include cardiac allograft rejection, kidney allograft rejection, liver allograft rejection, atherosclerosis, congestive heart failure, systemic lupus erythematosus,
                                                                                                                                                                                                                                                                                     rheumatoid arthritis, osteoarthritis or cytomegalovirus infection
                888888888888888
```

Sequence 50 BP; 13 A; 6 C; 16 G; 15 T; 0 U; 0 Other;

```
ó
                      Gaps
                      .
0
                                                                50
                                                        1 TGATGTTTGATGACCTATGAATCTATTTAGGGAGACACAGAGGTGGCTGGG
                                           3724 TGATGTTTGATGGACCTATGAATCTATTTAGGGAGACACAGATGGCTGGG
 Length 50;
                      0; Indels
Score 50; DB 1;
Pred. No. 0.15;
    100.0%; Preq. ....
 1.3%;
                       Conservative
           Local Similarity
                       50;
  Query Match
               Best Loca
Matches
                                             ò
                                                                  g
```

399Reverse primer for RFLP assay of human TLR4 gene. AAC84833 standard; DNA; 30 (first entry) 20-APR-2001 AAC84833; 

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; RFLP; ss.

Homo sapiens.

WO200077204-A1.

21-DEC-2000.

08-JUN-2000; 2000WO-US015723

99US-00329515 10-JUN-1999;

(CELL-) CELLULAR GENOMICS INC. 19-MAR-2001; 2001US-0276474P.

Velleca MA, Mellman I; WPI; 2002-759890/82.

(IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E.

Schwartz DA, Schutte BC; Lorenz E,

WPI; 2001-061872/07

Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll

Example 2; Page 42; 97pp; English.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, sepsis

```
ö
chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), sylvalomehitis, and adder disease, menumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. The present sequence represents a reverse primer used to detect a nucleotide change at codon 399 of the human TLR4 gene,
                                                                                                                                                                                                                                                                                                                                                                                                                             Human, TLR, Toll-like receptor; dendritic cell associated protein; autoimmune disorder; psoriasis; inflammatory bowel disease; asthma; multiple sclerosis; lupus erythematosus; rheumatoid arthritis; cancer; type I diabetes; infectious disease; gene therapy; immunosuppressive; antinflammatory; neuroprotective; dermatological; antibacterial; virucide; cytostatic; reverse transcription; RT; PCR; primer; ss.
                                                                                                                                                                                       Gaps
                                                                                                                                                                                       .;
0
                                                                                                                                                                                                                                                                                                                                                                                                       Human TLR4 (Toll-like receptor) DNA specific RT-PCR primer #2
                                                                                                     used in a PCR based RFLP assay to genotype patients for TLR4
                                                                                                                                                         / Match 0.8%; Score 30; DB 1; Length 30; Local Similarity 100.0%; Pred. No. 12; les 30; Conservative 0; Mismatches 0; Indels
                                                                                                                              Sequence 30 BP; 10 A; 4 C; 9 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                  1706 AGCALTTAACTCACTCTCCAGTCTTCAGGT 1735
                                                                                                                                                                                                                                    19-MAR-2002; 2002WO-US008122.
                                                                                                                                                                                                                                                                                                                      AAD46775 standard; DNA; 30
                                                                                                                                                                                                                                                                                                                                                                               (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              26-SEP-2002.
                                                                                                                                                                                                                                                                                                                                                                               27-JAN-2003
                                                                                                                                                                                                                                                                                                                                                  AAD46775;
                                                                                                                                                                 Query Match
                                                                                                                                                                                                                                                                                                            AAD46775/c
                                                                                                                                                                                            Matches
                                                                                                                                                                                                                                                                                                                                        88888888888888
                                                                                                                                                                                                                                                   셤
```

The invention relates to a method for generating a dendritic cell associated protein. The invention also relates to compositions and methods for generating an antibody against a dendritic cell associated protein. The methods and compositions are useful for diagnosing and treating diseases associated with altered dendritic cell activity such autoimmune disorders, e.g. psoriasis, inflammatory bowel disease, asthma, multiple sclerosis, lugus erychematosus, rheumatoid arthritis or type I diabetes, and cancer or infectious disease. The invention is also used in gene therapy. The present sequence is a RI (reverse transcription)-PCR perimer used for amplifying human TLM (Toll-like receptor) DNA. This sequence is used to illustrate the method of the invention Isolating dendritic cell associated protein using an agent which alters its expression or activity, useful in diagnosing and treating disorders with altered expression or activity of the protein, such as autoimmune Example 9; Page 56; 73pp; English. disease and cancer.

Sequence 30 BP; 9 A; 8 C; 10 G; 3 T; 0 U; 0 Other;

```
ö
                               Gaps
                              ;
0
        Length 30,
                              0; Indels
      DB 1;
0.8%; Scc...
100.0%; Pred. No. ...
0; Mismatches
                                                   2701 CCTCCTGAGGCATTTCTTGCCCAGCTGGGT 2730
                                                                  30 CCTCCTGAGGCATTTCTTGCCCAGCTGGGT
                 Best Local Similarity 100.
Matches 30; Conservative
         Query Match
                                                                   a
                                                   ઠે
```

AAC84832 standard; DNA; 31 BP

AAC84832;

(first entry) 20-APR-2001 399Forward primer for RFLP assay of human TLR4 gene.

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; RFLP; ss.

Homo sapiens.

WO200077204-A1

21-DEC-2000

08-JUN-2000; 2000WO-US015723

99US-00329515 10-JUN-1999;

(IOWA.) UNIV IOWA RES FOUND. (LORE/) LORENZ E.

Schutte BC; Lorenz E, Schwartz DA,

WPI; 2001-061872/07

Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNJ amplified from a biological sample encodes a portion of variant toll receptor

Claim 35; Page 42; 97pp; English.

methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with gram-negative sepsis, presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chromic airway disease, aethma, arthritis, local and systemic chronic airway disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic and cystic fibrosis. The present sequence represents a forward primer used to detect a nucleotide change at codon 399 of the human TLR4 gene, we have a primer and system in a PCR based RFLP assay to genotype patients for TLR4 The invention relates to human toll receptor 4 (TLR4) nucleic acid and 

Sequence 31 BP; 7 A; 3 C; 10 G; 11 T; 0 U; 0 Other;

0.8%; Score 29.4; DB 1; Length 31;

ö Gaps ö Indels 1, 0; Mismatches Pred. No. 14; 96.8%; 30; Conservative Best Local Similarity Matches 30; Conserv

ò ద RESULT 5 AAC84830

AAC84830 standard; DNA; 30

AAC84830;

(first entry) 20-APR-2001

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; RFLP; ss. 299Forward primer for RFLP assay of human TLR4 gene.

WO200077204-A1.

21-DEC-2000.

08-JUN-2000; 2000WO-US015723. 10-JUN-1999; 99US-00329515.

(IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E.

Schwartz DA, Schutte BC; Lorenz E,

WPI; 2001-061872/07.

altered innate immunity involves detecting or determining whether DN amplified from a biological sample encodes a portion of variant toll dentifying humans at risk of, or having indication associated with receptor 4.

Claim 33; Page 42; 97pp; English.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 unclaic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis. Creverity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for pre-term infants, agents which alter TLR4 activity are useful for pre-term infants, agents which alter TLR4 activity are useful for pre-term infants, agents which alter TLR4 activity are useful for in pre-term infants, agents which alter TLR4 activity are useful for creventing or ameliorating infection by gram-negative bacteria, sepsis induced chinded by gram-negative bacteria, negative sepsis induced chinded disease, preumonia, bronchitis, chronic syndrome (STRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, preumonia, bronchitis, chronic cobstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. The present sequence represents a forward primer used to detect a nucleotide change at codon 259 of the human TLR4 gene, and cystic fibrosis. used in a PCR based RFLP assay to genotype patients for TLR4

Sequence 30 BP; 9 A; 8 C; 4 G; 9 T; 0 U; 0 Other;

Gaps ô 0.7%; Score 28.4; DB 1; Length 30; 96.7%; Pred. No. 18; 1: Indel8 Query Match 0.7 Best Local Similarity 96.7 Matchés 29; Conservative

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 unclaic acid is useful as a diagnostic presence of TLR4 mutation is associated with pram-negative sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, sepsis induced by gram-negative bacteria, sepsis chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (STRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic and cystic fibrosis. The present sequence represents a reverse primer used to detect a nucleotide change at codon 259 of the human TLR4 gene, and cystic fibrosis.
                                                                                                                                                                                                                                                                                                                   TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; RFLP; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNJ amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          used in a PCR based RFLP assay to genotype patients for TLR4
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0.7%; Score 27; DB 1; Length 27;
                                                                                                                                                                                                                                                                             299Reverse primer for RFLP assay of human TLR4 gene.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 27 BP; 10 A; 8 C; 3 G; 6 T; 0 U; 0 Other;
1030 GATTAGCATACTTAGACTACTTACCTCGATG 1059
                           1 GATTAGCATACTTAGACTACTACCTCCATG 30
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Example 2; Page 42; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     10-JUN-1999; 99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (IOWA ) UNIV IOWA RES FOUND
                                                                                                                                                   AAC84831 standard; DNA; 27
                                                                                                                                                                                                                                      (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Lorenz E, Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens
                                                                                                                                                                                                                                      20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   receptor 4.
                                                                                                                                                                                              AAC84831;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    LORE/)
                                                                                                                            셤
    ઢ
```

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis.

The presence of TLR4 mutation is associated with gram-negative sepsis in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, included inflammatory disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (RDS), choolic obstructive pulmonary disease, pneumonia, bronchitis, chronic and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for
                                                                                                                                                                                                                                                            TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNI amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                              Human TLR4 gene exon 4 amplifying reverse primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        amplifying the exons of human TLR4 gene
Lorenz E, Schwartz DA, Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Example 1; Page 31; 97pp; English.
                                                                                                                 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          .0-JUN-1999; 99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (IOWA ) UNIV IOWA RES FOUND.
(LORE/) LORENZ B.
                                                                                                                 AAC84807 standard; DNA; 26
                                                                                                                                                                                          (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                           WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                       Homo sapiens.
                                                                                                                                                                                            20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         receptor 4.
                                                                                                                                                       AAC84807;
                   27
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (LORE/)
                                                                             RESULT 7
AAC84807/c
                                                                                                                   g
```

·

Gaps

; 0

0; Indels

100.0%; Pred. No. 21; tive 0; Mismatches

Local Similarity 100.0

Matches

Query Match

Sequence 26 BP; 8 A; 6 C; 2 G; 10 T; 0 U; 0 Other;

FirR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis;

respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyclonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

08-JUN-2000; 2000WO-US015723.

WO200077204-A1.

21-DEC-2000.

Ното варіепв

99US-00329515.

10-JUN-1999;

(IOWA ) UNIV IOWA RES FOUND.

(LORE/) LORENZ E.

1

```
The present invention describes an anti-CD14 antibody, which has a function of inhibiting the binding of CD14 to the Toll-like receptor (TLR). The anti-CD14 antibody can specifically recognise the epitope containing the domain from numbers 269-315 in human CD14 of the sequence in AAG68127 or a part of it. Anti-CD14 antibody has antibacterial, immunosuppressive, antipyretic, hypertensive, immunostimulant, haemostatic and vasotropic activities. The antibody together with other polypeptides are applicable in drugs for treating bacterial infection as well as sepais, fever, hypotension, leukopaenia, thrombopaenia, shock and multi-organ failure. AAG68127 to AAG68137 and AAI71210 to AAI71295
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Anti-CD14 antibody or its fragment inhibiting the binding of CD14 to Toll -like receptor, applicable in drugs for treating bacterial infection as well as sepsis, fever, hypotension, leukopenia, thrombopenia and shock..
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               represent sequences used in the exemplification of the present invention
                                                                                                                                                                                     Human; Toll like receptor; TLR; CD14; antibody; anti-CD14 antibody; TLK/CD14 binding inhibitor; antibacterial; immunosuppressive; antipyretic; hypertensive; immunostimulant; haemostatic; vasotropic; bacterial infection; sepsis; fever; hypotension; leukopaenia; thrombopaenia; shock; multi-organ failure; ss.
                                                                                                                                                 Human Toll like receptor 4 PCR antisense primer 1 SEQ ID NO:9.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DB 1; Length 24;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Seguence 24 BP; 11 A; 5 C; 3 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Takahashi
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Score 24;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Example 2; Page 170; 202pp; Japanese.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Shirakawa K,
                                 AAI71236 standard; DNA; 24 BP
                                                                                                                                                                                                                                                                                                                                                                                                                        02-APR-2001; 2001WO-JP002869.
                                                                                                                                                                                                                                                                                                                                                                                                                                                          31-MAR-2000; 2000JP-00099617.
22-NOV-2000; 2000JP-00356719.
28-MAR-2001; 2001US-00906158.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (MOCH ) MOCHIDA PHARM CO LTD
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0.6%;
                                                                                                           23-JAN-2002 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Furusako S, Mori S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2001-616487/71.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                            WO200172993-A1
                                                                                                                                                                                                                                                                                                         sapiens
                                                                                                                                                                                                                                                                                                                                                                                04-OCT-2001
                                                                       AAI71236;
               AAI71236/c
ID AAI712
                                                                                                                                                                                                                                                                                                       Ношо
RESULT 8
```

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis.

The presence of TLR4 mutation is associated with gram-negative sepsis in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LDS (lipopolysaccharide) induced chronic airway disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory response inflammatory disease, asthma, arthritis, local and systemic inflammatory disease, asthma, arthritis, local and systemic syndrome (SIRS) or acute respiratory distress syndrome (ARS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS;
                                                                                                                                                                                                                                                                                                                                                                                                                                      Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNJ amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Score 24; DB 1; Length 24;
Pred. No. 37;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Seguence 24 BP; 6 A; 5 C; 4 G; 9 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human TLR4 gene exon 4 amplifying reverse primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                859
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  24 AGAAATTAGGCTTCATAAGCTGAC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                836 AGAAATTAGGCTTCATAAGCTGAC
                                                                                                                                                                                                                                                                                                                                                                     Schutte
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.6%; 8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAC84810 standard; DNA; 24
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           24; Conservative
                                                                                                                                                                                                                                                                                                                                                                   Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 receptor 4.
                                                                                                                                                                                                                                                                                                                                                                   Corenz E,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AAC84810;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Local
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RESULT 10
```

ö

Gaps

; 0

0; Indels

100.0%; Pred. No. 37; ive 0; Mismatches

24; Conservative

Matches

ð

1137 GACTTTTCTTATAATTTCGGATGG 1160

GACTTTTCTTATAATTTCGGATGG 1

AAC84805 standard; DNA; 24 BP.

RESULT 9

Human TLR4 gene exon 4 amplifying reverse primer.

20-APR-2001 (first entry)

AAC84805;

AAC84805/c
ID AAC848
XX
AC AAC848
XX
XX
XX
DF 20-APR
DE Human

ω

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 mucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 muctation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for conversity of smellocating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, sepsis cinduced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic conditions such as systematic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, nocal gram-negative bacterial infection and cystic fibrosis. Sequences AACM34776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human, TLR; Toll-like receptor; dendritic cell associated protein; autoimmune disorder; psoriasis; inflammatory bowel disease; asthma; multiple sclerosis; lupus erythematosus; rheumatoid arthritis; cancer; type I diabetes; infectious disease; gene therapy; immunosuppressive; antiinflammatory; neuroprotective; dermatological; antibacterial;
chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pydonosphritis; bronchitis; acute respiratory distress some; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                              Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human TLR4 (Toll-like receptor) DNA specific RT-PCR primer #1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Score 24; DB 1; Length 24; Pred. No. 37;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 24 BP; 8 A; 5 C; 2 G; 9 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     0.6%; Scc...
100.0%; Pred. No. 3...
... 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1464 TTGAAACAAATGAGTGAGTTTTCA 1487
                                                                                                                                                                                                                                                                                                                                                                    Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        rrgaaacaaargagrgagrrrrca 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             BP.
                                                                                                                                                                                                                           08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                 99US-00329515.
                                                                                                                                                                                                                                                                                                     (IOWA ) UNIV IOWA RES FOUND.
(LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAD46774 standard; DNA; 24
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
Best Local Similarity 100.1
Matches 24, Conservative
                                                                                                                                                                                                                                                                                                                                                                      Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2001-061872/07.
                                                                                                                                             WO200077204-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            27-JAN-2003
                                                                                                                                                                                                                                                                   10-JUN-1999;
                                                                                                       Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               receptor 4.
                                                                                                                                                                                   21-DEC-2000
                                                                                                                                                                                                                                                                                                                                                                      Lorenz E,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        24
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAD46774;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     RESULT 11
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      8
```

```
The invention relates to a method for generating a dendritic cell associated protein. The invention also relates to compositions and methods for generating an antibody against a dendritic cell associated protein. The methods and compositions are useful for diagnosing and treating diseases associated with altered dendritic cell activity such as autoimmune disorders, e.g. psoriasis, inflammatory bowel disease, asthma, multiple sclerosis, lupus erythematosus, rheumatory bowel disease, type diabetes, and cancer or infectious disease. The invention is also used in gene therapy. The present sequence is a RT (reverse transcription)-PCR primer used for amplifying human TLR (Toll-like receptor) DNA. This
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Isolating dendritic cell associated protein using an agent which alters its expression or activity, useful in diagnosing and treating disorders with altered expression or activity of the protein, such as autoimmune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Immunosuppressive; antibacterial; anti-CD14 antibody; epitope; sepsis;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
virucide; cytostatic; reverse transcription; RT; PCR; primer; 88
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         sequence is used to illustrate the method of the invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.6%; Score 24; DB 1; Length 24;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Anti-CD14 monoclonal antibody related oligonucleotide #2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 24 BP; 4 A; 6 C; 7 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Pred. No. 37;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                100.0%; Prec. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      2442 CTGAGCAGTCGTGCTGGTATCATC 2465
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 24
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              creascastcstscrestarcarc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Example 9; Page 56; 73pp; English.
                                                                                                                                                                                                                                                                                                                                                                                      (CELL-) CELLULAR GENOMICS INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      28-SEP-2001; 2001WO-JP008563.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 22-NOV-2000; 2000JP-00356719.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (MOCH ) MOCHIDA PHARM CO LTD.
                                                                                                                                                                                                                                                         19-MAR-2002; 2002WO-US008122.
                                                                                                                                                                                                                                                                                                                        19-MAR-2001; 2001US-0276474P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AAL41014 standard; DNA; 24
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              24; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                 Velleca MA, Mellman I;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 2002-759890/82
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          disease and cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   human CD14; ds
                                                                                                                                   WO200274921-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WO200242333-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Unidentified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        11-OCT-2002
                                                                        Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           30-MAY-2002.
                                                                                                                                                                                              26-SEP-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AAL41014;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Н
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Best Loc
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 12
AAL41014/C
ID AAL41
XX
AC AAL41
XX
XX
DT 11-0C
XX
XX
DE ANTi-
XX
XM
IMMUN
XM
IMMUN
XM
IMMUN
XM
IMMON

         ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 g
```

Mori

```
Anti-CD14 monoclonal antibody which inhibits CD14/T lymphocyte receptor binding by specifically recognizing epitope in human CD14 domain to prevent interaction and suppress cell activation, useful for treating
 Shirakawa K.
                  WPI; 2002-454920/48.
 Furusako S,
                                                                    вервів.
```

The invention relates to an anti-CD14 antibody which can specifically recognise an epitope containing a part of a domain with not less than 8 amino acids in human CD14 in the region from positions 269-315 in a ful defined sequence of 356 amino acids as given in the specification. The antibody is useful in drug compositions for treating sepsis and for screening remedies for sepsis. This polynucleotide sequence represents anti-CD14 related oligonacleotide of the invention 0.6%; Score 24; DB 1; Length 24; 00.0%; Pred. No. 37; Sequence 24 BP; 11 A; 5 C; 3 G; 5 T; 0 U; 0 Other; Example 2; Page 45; 156pp; Japanese. Query Match

100.0%; Pred. No. 37; cive 0; Mismatches 1137 GACTTTTCTTATAATTTCGGATGG 1160 24 GACTTTTCTTATAATTTCGGATGG 1 24; Conservative ò 원

Local Similarity

Best Loca Matches

ACC43816

ACC43816 Btandard; DNA; 24

ACC43816;

(first entry) 11-AUG-2003

Sense PCR primer for human toll-like receptor 4 (TLR4) cDNA.

Toll-like receptor; TLR; central nervous system; CNS; neurodegenerative disorder; Alzheimer's disease; Parkinson's disease; Pick's disease; multiple sclerosis; stroke; PCR; primer; ss.

Homo sapiens

EP1288226-A1

05-MAR-2003

03-SEP-2001; 2001EP-00203325

03-SEP-2001; 2001EP-00203325

(NEDE ) NEDERLANDSE ORG TOEGEPAST.

Van Noort JM

WPI; 2003-344752/33.

Modifying the expression of Toll-like receptors (TLR), useful for influencing neurodegeneration or neuroprotection in the human CNS, comprises contacting CNS cells with a TLR-expression modifying agent, e.g. alpha B-crystallin.

Disclosure; Page 7; 20pp; English.

The specification describes a method of modifying the expression of at toll-like receptor (TLR) in cells of the human central nervous system (CNS). The method comprises contacting the cells with a TLR-expression modifying agent, selected from substances that are endogenous to the human CNS and its parts or variants, that is capable of altering the expression of a TLR in the cells. The TLR-expression modifying agent is.

inhibiting a neurodegenerative process and/or stimulating a neurodegenerative neuroprotective process in a human being afflicted by a neurodegenerative disorder such as Alzheimer's disease, Parkinson's disease, Pick's disease, multiple sclerosis or stroke. The present PCR primer was used to amplify cDNA encoding human TERI. The primer was used to determine the amount of TLR present in human glia cells in a semi-quantitative RT-PCR pharmaceutical composition for retarding or useful for preparing a 888888888

Sequence 24 BP; 8 A; 2 C; 10 G; 4 T; 0 U; 0 Other;

Gaps ö Score 24; DB 1; Length 24; Pred. No. 37; 0; Indels 100.0%; Pred. No. 3,, tive 0; Mismatches 0.68; 24; Conservative Local Similarity Query Match Matches

ö

· 2218 AGGACTGGGTAAGGAATGAGCTAG 2241 24 1 AGGACTGGGTAAGGAATGAGCTAG ઠે 셤

ADR83260

BP ADR83260 standard; RNA; 24 ADR83260;

ö

Gaps ö

0; Indels

Human toll-like receptor 4 miRNA target region.

(first entry)

02-DEC-2004

immune disease; nerve disorder; amyotrophic lateral sclerosis; Parkinson's disease; Alzheimer's disease; inflammatory disease; siRNA silencing precursor; cytostatic; immunosuppressive; nootropic; neuroprotective; antiinflammatory; immunotherapy; toll-like receptor 4. human; 88; miRNA; microRNA; ontogenesis; cell therapy; cancer;

Homo sapiens

WO2004076622-A2.

10-SEP-2004.

10-FEB-2004; 2004WO-JP001433

10-FEB-2003; 2003US-0445829P

(NAAD-) NAT INST ADVANCED IND SCI & TECHNOLOGY.

Taira K, Kawasaki H;

WPI; 2004-653393/63.

Modulating expression of a target gene in a cell, for treating cancer, an immune disease, or a nerve disorder, comprises introducing into the cell a polynucleotide that forms a duplex region with an mRNA transcribed from target gene 

Claim 2; SEQ ID NO 162; 865pp; English

This invention relates to a novel method for modulating the expression of a target gene in a cell. Specifically, it refers to the introduction into a call of a polynocleoride that forms a duplex region with an mRNA transcribed from the target gene, where the duplex region comprises a mammalian miRNA target region i.e. a non-coding microRNA (miRNA) that regulates mRNA at a post-transcriptional level. The present invention describes a method for controlling ontogenesis of a mammal, function of a mammalian cell, differentiation of a mammalian cell, differentiation of a mammalian cell, differentiation of a mammalian cell or viability of a mammalian cell in the post-transcriptional phase, which comprises introducing a plasmid vector comprising a promoter and nucleic acid molecule expressing an miRNA or sirNA silencing precursor to the miRNA.

Cocordingly, it provides a cell therapy method for treating cancer, immune disease, nerve disorder (e.g. amyotrophic lateral sclerosis, immune disease, or Alzheimer's disease) or an inflammatory disease by introducing into the cell the miRNA, siRNA silencing precursor to the

ô

```
methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 mucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, is preventing or ameliorating infection by gram-negative bacteria, sepsis chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS) syndrome (SIRS) or acute respiratory distress syndrome (ARDS) systematic and systematic phenomery disease, local gram-negative bacterial infection obstructive pulmonary disease, local gram-negative bacterial infection
                pharmaceutical compositions that exhibit cytostatic, immunosuppressive, norotropic, neuroprotective and antiinflammatory activities and hence can be used for immunotherapy. This oligonucleotide sequence is a human miRNA target region derived from a target gene of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            invention relates to human toll receptor 4 (TLR4) nucleic acid and
                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DN amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                    ö
miRNA or the plasmid vector. As such, they can be developed
                                                                                                                                                        Score 24; DB 1; Length 24;
Pred. No. 37;
8; Mismatches 0; Indels
                                                                                                                    Sequence 24 BP; 6 A; 7 C; 3 G; 0 T; 8 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                            2278 TCTGCCTTCACTACAGAGACTTTA 2301
                                                                                                                                                                                                                                                                 1 UCUGCCUUCACUACAGAGACUUUA 24
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Schutte
                                                                                                                                                                                                                                                                                                                                                                                      ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        99US-00329515.
                                                                                                                                                              0.6%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                    AAC84796 standard; DNA; 23
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20-APR-2001 (first entry)
                                                                                                                                                                                                      16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 2001-061872/07
                                                                                                                                                                                 Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          21-DEC-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     receptor 4.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ŭ
                                                                                                                                                                                                                                                                                                                                                                                                                          AAC84796;
                                                                                                                                                              Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Lorenz
                                                                                                                                                                                                        Matches
                                                                                                                                                                                                                                                                                                                                                                     AAC8479
                                                                                                                                                                                                                                                                                                                                                                                        8888888
                                                                                                                                                                                                                                                8
                                                                                                                                                                                                                                                                                요
```

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered individuals immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, induced by gram-negative bacteria, induced chronic airway disease conditions such as systematic inflammatory response inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS).

Syndrome (SIRS) or acute respiratory distress syndrome (ARDS).

Cobstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                          Gaps
                                                                                                                                       ö
                                                                                                   1; Length 23;
                                                                                                                                       0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                  Human TLR4 gene exon 4 amplifying forward primer.
                                                            Sequence 23 BP; 5 A; 6 C; 3 G; 9 T; 0 U; 0 Other;
                                                                                                     DB 1
                                                                                                                                            0; Mismatches
                                                                                                     Score 23;
Pred. No.
                                                                                                                                                                              2459 TATCATCTTCATTGTCCTGCAGA 2481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                BC;
                                                                                                                                                                                                    1 TATCATCTTCATTGTCCTGCAGA 23
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Schwartz DA, Schutte
                                                                                                                                                                                                                                                                                                                    ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    99US-00329515.
                                                                                                                       100.08;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (IOWA ) UNIV IOWA RES FOUND.
                                                                                                       0.68;
                                                                                                                                                                                                                                                                                                                    AAC84783 standard; DNA; 23
                                                                                                                                                                                                                                                                                                                                                                                               (first entry)
                                                                                                                       Similarity 100.
23; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                                               20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        receptor 4.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Lorenz E,
                                                                                                                                                                                                                                                                                                                                                             AAC84783;
                                                                                                       Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (LORE/)
                                                                                                                           Local
                                                                                                                                            Matches
                                                                                                                                                                                                                                                                                 RESULT 16
                                                                                                                                                                                                                                                                                                      AAC84783
                                                                                                                                                                                                                                                                                                                      ន្តដូន
```

11

ö

Gaps

; 0

```
TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis;
                                                                                                                                                                                                                                                                                respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                       DB 1; Length 23; 44;
                                                 0; Indels
Sequence 23 BP; 10 A; 0 C; 8 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                           Human TLR4 gene exon 4 amplifying forward primer.
                                                  Mismatches
                     Query Match 0.6%; Score 23; Best Local Similarity 100.0%; Pred. No. Matches 23; Conservative 0; Mismatc
                                                                         943 TGGGAGAATTTAGAAATGAAGGA 965
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ВС;
                                                                                       1 TGGGAGAATTTAGAAATGAAGGA 23
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Schutte
                                                                                                                                                               AAC84787 standard; DNA; 23 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                     08-JUN-2000; 2000WO-US015723
                                                                                                                                                                                                                                                                                                                                                                                                                                                             99US-00329515
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (IOWA ) UNIV IOWA RES FOUND.
                                                                                                                                                                                                                20-APR-2001 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Lorenz E, Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                    WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                             10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                            Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                            21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     receptor 4.
                                                                                                                                                                                          AAC84787;
```

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory disease used as systematic inflammatory disease, local gram-negative bacterial infection obstructive pulmonary disease, local gram-negative bacterial infection costic the exons of human TLR4 gene

```
Sequence 23 BP; 7 A; 5 C; 4 G; 7 T; 0 U; 0 Other;
```

0.6%; Score 23; DB 1; Length 23;

Query Match

ö

Gaps

ö

0; Indels

0.6%; Score 23; DB 1; Length 23; 00.0%; Pred. No. 44;

100.0%; Pred. ....

Query Match Best Local Similarity 100. Matches 23; Conservative

Sequence 23 BP; 6 A; 1 C; 8 G; 8 T; 0 U; 0 Other;

```
methods to identify polymorphisms at the human Tike 10 cus and to identify methods to identify polymorphisms at the human Tike 10 cus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant Tike nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human Tike gene. Since the presence of Tike mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter Tike activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, included chronic airway disease conditions such as systemic inflammatory disease. Sequences and sease, porchiais, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for
                     ö
                                                                                                                                                                                                                                                                                                                                                                        TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               The invention relates to human toll receptor 4 (TLR4) nucleic acid and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Identifying humans at risk of, or having indication associated with
                     ö
                     Indels
                     .;
0
                                                                                                                                                                                                                                                                                                                                   Human TLR4 gene exon 4 amplifying reverse primer.
  44:
                     0; Mismatches
100.0%; Pred. No.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                and cystic fibrosis. Sequences AACG4''00 amplifying the exons of human TLR4 gene
                                                            1410 TCAAACTTCTTGGGCTTAGAACA 1432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Schwartz DA, Schutte BC;
                                                                                                   1 rchanctrcrredecrradadda 23
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           10-JUN-1999; 99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (IOWA ) UNIV IOWA RES FOUND (LORE/) LORENZ E.
                                                                                                                                                                                                       AAC84808 standard; DNA; 23
                                                                                                                                                                                                                                                                                         20-APR-2001 (first entry)
                   23; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2001-061872/07.
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 receptor 4.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Lorenz E,
                                                                                                                                                                                                                                                AAC84808;
                                                                                                                                                                  RESULT 18
AAC84808/c
                   Matches
                                                                                                                                                                                                                               ð
```

RESULT 19

셤 ð

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered inmunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, pre-term delivery and respiratory distress syndrome preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, local and systemic circlammatory disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (STRS) or acute respiratory distress syndrome (RTRS).

Syndrome (STRS) or acute respiratory distress syndrome (RDRS), chounce obstructive pulmonary disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, ARCB4776-823 represent PCR primers for and cystic fibrosis. Sequences.
                                                                                TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Score 22; DB 1; Length 22;
Pred. No. 52;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human TLR4 gene exon 4 amplifying reverse primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 22 BP; 6 A; 4 C; 4 G; 8 T; 0 U; 0 Other;
                                         Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               and cystic fibrosis. Sequences AAC84776·
amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1323 TTCAAAGGTTGCTGTTCTCAAA 1344
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Lorenz E, Schwartz DA, Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1 rrcaaaggrigcrgricrcaaa 22
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0.00,
100.0%; Pre
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      BP
                                                                                                                                                                                                                                                                                                                                                                                                    08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.6%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            IOWA RES FOUND
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AAC84806 standard; DNA; 22
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (first entry)
20-APR-2001 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Local Similarity 100.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                         WO200077204-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (IOWA ) UNIV
                                                                                                                                                                                                                                                                                                                                                                                                                                                10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 20-APR-2001
                                                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                                                                                                                   21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    receptor 4.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AAC84806;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAC84806/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           BXHXHXB
  ద
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        The specification describes a method of modifying the expression of at a toll—like receptor (TLR) in cells of the human central nervous system (CNS). The method comprises contacting the cells with a TLR-expression modifying agent, selected from substances that are endogenous to the human CNS and its parts or variants, that is capable of altering the expression of a TLR in the cells. The TLR-expression modifying agent is useful for preparing a pharmaceutical composition for retarding or inhibiting a neurodegenerative process and/or stimulating a neurodegenerative process and/or stimulating a neurodegenerative disease, parkinson is disease, pick's disease, multiple sclerosis or stroke. The present PCR primer was used to amplify CDNA encoding human TLRI. The primer was used to determine the amount of TLR present in human glia cells in a semi-quantitative RT-PCR
                                                                                                                                                                                                                                                                                                                                                          Toll-like receptor; TLR; central nervous system; CNS; neurodegenerative disorder; Alzheimer's disease; Parkinson's disease; Pick's disease; multiple sclerosis; stroke; PCR; primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Modifying the expression of Toll-like receptors (TLR), useful for influencing neurodegeneration or neuroprotection in the human CNS, comprises contacting CNS cells with a TLR-expression modifying agent, e.g. alpha B-crystallin.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                              Antisense PCR primer for human toll-like receptor 4 (TLR4) cDNA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ch 0.6%; Score 22.4; DB 1; Length 24; 1. Similarity 95.8%; Pred. No. 55; 23; Conservative 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 24 BP; 3 A; 8 C; 5 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      2627 GAATCCAGAAGGAACAGTGGGTAC 2650
         CCCACATTGAAACTCAAATCTCT 1222
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      CCCACATTGAAACTCAAATCTCT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (NEDE ) NEDERLANDSE ORG TOEGEPAST.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Disclosure; Page 7; 20pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                03-SEP-2001; 2001EP-00203325.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            03-SEP-2001; 2001EP-00203325.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AAC84786 standard; DNA; 22
                                                                                                                                                                        ACC43826 standard; DNA; 24
                                                                                                                                                                                                                                                                11-AUG-2003 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2003-344752/33.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Van Noort JM;
                                                                                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        EP1288226-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      05-MAR-2003.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAC84786;
         1200
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    24
                                                                                                                                                                                                                         ACC43826;
                                                        23
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Local
```

ö

Gaps

; 0

0; Indels

Best Loc Matches

8 원 RESULT 20

AAC84786 ID AAC8 XX AC AAC8 XX

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 uncleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, pre-term delivery and respiratory disteress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, sepsis induced by gram-negative bacteria, tDR (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AGC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Vascular endothelial growth factor receptor; VEGF receptor; flt-1; flk-1; KDR; hammerhead ribozyme; hairpin ribozyme; cleavage;
                  TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Mouse flt-1 VEGF receptor hammerhead ribozyme #241.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 22 BP; 7 A; 7 C; 4 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   TCTAGAGGGCCTGTGCAATTTG 1013
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        22 TCTAGAGGGCCTGTGCAATTTG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AAX73769 standard; RNA; 27 BP
                                                                                                                                                                                                                                                                               08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                      99US-00329515.
                                                                                                                                                                                                                                                                                                                                                            (IOWA ) UNIV IOWA RES FOUND
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                          Lorenz E, Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2001-061872/07
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Similarity
                                                                                                                                                                                                                                                                                                                                                                                  (LORE/) LORENZ E.
                                                                                                                                                                                                WO200077204-A1.
                                                                                                                                                           Homo sapiens.
                                                                                                                                                                                                                                                                                                                    10-JUN-1999;
                                                                                                                                                                                                                                        21-DEC-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         28-JUL-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          22;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   992
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     receptor
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AAX73769
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Local
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Matches
ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   원
```

Schutte BC;

```
Nucleic acid molecule modulating VEGF receptor(s) gene expression or mRNA stability - useful for treating e.g. tumour angiogenesis, psoriasis, rheumatoid arthritis, etc., in a human patient.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          synthesis, expression and/or stability of a mRNA encoding 1 or more receptors of vascular endothelial growth factor (VEGF). A patient (preferably human) having a growth factor (VEGF). A patient fms-like tyrosine kinase 1 (flt-1), kinase insert domain containing arceptor (KDR) and/or foetal liver kinase 1 (flk-1) (e.g. tumour anglogenesis, ocular diseases, psoriasis and rheumatoid arthritis) car treated by administering the nucleic acid molecule or the expression
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Vascular endothelial growth factor receptor; VEGF receptor; flt-1; flk-1; KDR; hammerhead ribozyme; hairpin ribozyme; cleavage; tumour angiogenesis; psoriasis; rheumatoid arthritis; ocular disease; fms-1ike tyrosine kinase 1; kinase insert domain containing receptor; foetal liver kinase 1; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                              The present invention describes nucleic acid molecules which modulate the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        vector to the patient. AAX67275 to AAX75752 represent specific examples of nucleic acid molecules from the present invention
              fms-like tyrosine kinase 1; kinase insert domain containing receptor;
foetal liver kinase 1; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gaps
ocular disease;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DB 1; Length 27;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      4; Indels
tumour anglogenesis; psoriasis; rheumatoid arthritis;
                                                                                                                                                                                                                                                                                                      Pavco P, Mcswiggen J, Stinchcomb D, Escobedo J;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 27 BP; 12 A; 4 C; 4 G; 0 T; 6 U; 1 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human flt1 VEGF receptor hammerhead ribozyme #309.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    5; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    2673 ACATCTATCTGAAGAGGAAAAATAAAA 2699
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Score 21.2;
Pred. No. 91
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1 ACAUCUGUCUGAUGANGAAACAUAAAA 27
                                                                                                                                                                                                                                                                                                                                                                                                                               Claim 9; Page 162; 218pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ВЪ
                                                                                                                                                                         96WO-US017480.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.6%;
                                                                                                                                                                                                      95US-0005974P.
                                                                                                                                                                                                                                                   (RIBO-) RIBOZYME PHARM INC. (CHIR.) CHIRON CORP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AAX67583 standard; RNA; 27
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    18; Conservative
                                                                                                                                                                                                                                                                                                                                WPI; 1997-259017/23
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Similarity
                                                                                                                                                                       25-OCT-1996;
                                                                                                         WO9715662-A2
                                                                                                                                                                                                                     11-JAN-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               28-JUL-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WO9715662-A2
                                                                                                                                        01-MAY-1997.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      01-MAY-1997
                                                            Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
Best Local S
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AAX67583;
                                                                              Мив вр.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 23
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAX67583
 ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
```

.. 0

Gaps ö

DB 1; Length 22; 52; 0; Indels

100.0%; Pred. ... Score 22; Pred. No.

0.6%;

```
synthesis, expression and/or stability of a mRNA encoding 1 or more receptors of vascular endothelial growth factor (VBGF). A patient (preferably human) having a condition associated with the level of the fms-like tyrosine kinase 1 (fll-1), kinase insert domain containing receptor (KDR) and/or foetal liver kinase 1 (flk-1) (e.g. tumour angiogenesis, ocular diseases, psoriasis and rheumatoid arthritis) can be treated by administering the nucleic acid molecule or the expression vector to the patient. AAX67275 to AAX7572 represent specific examples of nucleic acid molecules from the present invention
                                                                                                                                                                                                       Nucleic acid molecule modulating VEGF receptor(s) gene expression or mRNA stability - useful for treating e.g. tumour angiogenesis, psoriasis, rheumatoid arthritis, etc., in a human patient.
                                                                                                                                                                                                                                                                                                       present invention describes nucleic acid molecules which modulate the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 27 BP; 12 A; 3 C; 4 G; 0 T; 7 U; 1 Other;
                                                                                                                                           Stinchcomb D, Escobedo J,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Score 21.2; I
Pred. No. 91;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    5; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    2673 ACATCTATCTGAAGAGGAAAATAAAA 2699
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Claim 9; Page 56; 218pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0.6%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          08-JUN-2000; 2000WO-US015723
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        99US-00329515
               96WO-US017480
                                                              96US-00584040
                                               95US-0005974P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (IOWA ) UNIV IOWA RES FOUND (LORE/) LORENZ E.
                                                                                              (RIBO-) RIBOZYME PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AAC84784 standard; DNA; 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20-APR-2001 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      18; Conservative
                                                                                                                                           Pavco P, Mcswiggen J,
                                                                                                             (CHIR ) CHIRON CORP
                                                                                                                                                                         WPI; 1997-259017/23
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WO200077204-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        10-JUN-1999;
                25-OCT-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Ното варіепв
                                               26-OCT-1995;
                                                               11-JAN-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAC84784;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Best_Loca
Matches
                                                                                                                                                                                                                                                                                                         The
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 24
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                셤
```

Lorenz E, Schwartz DA, Schutte BC;

```
The intention relates to numeral coil receptor; (Link) indicate and to identify polymorphisms at the human Tik4 locus and to identify individuals at risk of, or having, an indication associated with altered inmunity. A variant Tik4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human Tik4 gene. Since the presence of Tik4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome seventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, ins (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (SIRS) or acute respiratory distress syndrome (AIRS), chronic obstructive pullmonary disease, neumonia, bronchitis, chronic obstructive pullmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequence AAC84776-823 represent PCR primers for
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ANDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                The invention relates to human toll receptor 4 (TLR4) nucleic acid and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                         altered innate immunity involves detecting or determining whether DN amplified from a biological sample encodes a portion of variant toll
                                      Identifying humans at risk of, or having indication associated with
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Identifying humans at risk of, or having indication associated with
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.6%; Score 21; DB 1; Length 21;
.00.0%; Pred. No. 62;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence 21 BP; 2 A; 5 C; 3 G; 11 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human TLR4 gene exon 4 amplifying reverse primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Schwartz DA, Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1093 TTTCTTCATTTTCCCTGGTGA 1113
                                                                                                                                         Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TTTCTTCATTTTCCCTGGTGA 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             100.08;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     08-JUN-2000; 2000WO-US015723
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           10-JUN-1999; 99US-00329515
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (IOWA ) UNIV IOWA RES FOUND.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AAC84803 standard; DNA; 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   21; Conservative
WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WPI; 2001-061872/07
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO200077204-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Lorenz E,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAC84803;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               RESULT 25
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AAC84803/
셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Š
```

ö

Gaps

; 0

4;

DB 1; Length 27; Indels

WO200077204-A1 10-JUN-1999; Homo sapiens. receptor 4. AAC84823; Query Match Best Local ( ઠે 유

altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll

Example 1; Page 31; 97pp; English.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 uncleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, lbS (llpopolysaccharide) induced chy gram-negative bacteria, LBS (llpopolysaccharide) induced chy gram-negative bacteria, lbS (llpopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic cobstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences ARCH776-821 represent PCR primers for amplifying the exons of human TLR4 gene

Sequence 21 BP; 7 A; 4 C; 5 G; 5 T; 0 U; 0 Other;

ö Gaps ö 0.6%; Score 21; DB 1; Length 21; 00.0%; Pred. No. 62; 0; Indels 100.0%; Pred. ... Matches 21; Conservative Similarity

AAC84823 standard; DNA; 21 BP

20-APR-2001 (first entry)

Human TLR4 gene exon 4 amplifying reverse primer.

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

21-DEC-2000

08-JUN-2000; 2000WO-US015723.

(IOWA ) UNIV IOWA RES FOUND (LORE/) LORENZ E. Lorenz E, Schwartz DA, Schutte BC;

WPI; 2001-061872/07.

Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll receptor 4.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LBS (lipopolygaccharide) induced chronic airway disease conditions such as systematic inflammatory response inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (SIRS) and cystic fibrosis. Sequences AAC64776-823 represent PCR primers for Gaps ö 0.6%; Score 21; DB 1; Length 21; 100.0%; Pred. No. 62; Astive 0; Mismatches 0; Indels Sequence 21 BP; 5 A; 6 C; 4 G; 6 T; 0 U; 0 Other; amplifying the exons of human TLR4 gene 2813 AGATATGCAGGCTGCTAATC 2833 Example 1; Page 31; 97pp; English. 21 AGATATGCAGGCTGCTAATC 1 21; Conservative Local Similarity Query Match Best Loca Matches g

ö

ACC70796/c RESULT 27

ACC70796 Btandard; DNA; 21

ВР

(first entry) ACC70796;

20-NOV-2003

Human Toll-like receptor 4, Tlr-4, PCR primer #2.

Human, PCR, primer; vulnerary; anti-tumour; antirheumatic; antiarthritic; antiarteriosclerotic; cytostatic; neointima; scar; plaque; blood vessel; Toll-like receptor 4; adventitial cell; Tlr-4; ss.

EP1302206-A1

16-APR-2003.

11-OCT-2001; 2001EP-00203846. 

11-OCT-2001; 2001EP-00203846.

(UYUT-) UNIV UTRECHT MEDISCH CENT. (UYUT-) RIJKSUNIV UTRECHT. Pasterkamp G; De Kleijn DPV,

WPI; 2003-484923/46.

Interfering with the formation of a neointima/scar and/or a plaque in a blood vessel, useful for modulating tumor growth, comprises providing a ligand capable of modulating Toll-like receptor activity of adventitial

Disclosure; Page 7; 23pp; English.

The present invention relates to a method for interfering with the formation of a neointima/scar and/or a plaque in a blood vessel by providing a ligand capable of modulating Toll-like receptor activity of

10001863-3.sl.rng

16

```
RESULT 30
                                                                                                                                                                                                                                                                                                                                                  X S
                                                                                                                                                                                                                    g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      엄
                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  The invention comprises therapeutic agents for preventing or treating diseases associated with endothelial disorders, the agents contain a 3-mino-1,2-benzisothiazole compound. The therapeutic agents of the invention are useful for preventing/treating: diseases associated with haemorthagic shock, alcohol induced cirrhosis, adult respiratory distress syndrome, chronic rheumatoid arthritis, ulcerative gastritis, Crohn's disease, glomerulonephritis, infectious carditis, systemic lupus erythematosus, scleroderma, Sjoegren's syndrome, multiple organ failure sequence represents a PCR primer that was used in the exemplification of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Agents for treating diseases associated with endothelial disorders, toxemia or toll like receptor signaling comprise new or known 3-amino-1,2-benzisothiazole compounds.
                  neointima/scar and/or a plaque in a blood vessel after stenting, angioplasty, heart transplantation, by pass surgery, arteriovenous shunting and infection, especially bacterial infection. The method is also useful for modulating tumour growth, and for modulating the effects of rheumatoid arthritis. The present sequence is a PCR primer for human
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             therapeutic agent; endothelial disorder; 3-amino-1; 2-benzisothiazole compound; endothelial disorders; toxaemia; severe toxaemia; toxic shock; haemorrhagic shock; alcock; alcohol induced cirrhosis; adult respiratory distress syndrome; chronic rheumatoid arthritis; ulcerative gastritis; Crohn's disease; glomerulonephritis; infectious carditis; systemic lupus erythematosus; ecleroderma; Sjoegren's syndrome; multiple organ failure; autoimmune disease; multiple sclerosis; PCR; primer; ss; human; toll-like receptor 4; TLR4.
The method is useful for reducing the formation of
                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                 ..
0
                                                                                                                                                                                                                                             0.6%; Score 21; DB 1; Length 21;
100.0%; Pred. No. 62;
iive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Human toll-like receptor 4 (TLR4) gene PCR primer #3
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Mori
                                                                                                                                                                                                 Seguence 21 BP; 8 A; 9 C; 0 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Nakamura M, Mizuno M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Example 4; SEQ ID NO 6; 180pp; Japanese.
                                                                                                                                                                                                                                                                                                                                                2376
                                                                                                                                                                                                                                                                                                                                                  2356 GAAAGGTGATTGTTGTGTGT
                                                                                                                                                                                                                                                                                                                                                                                              21 GAAGGTGATTGTTGTGTGT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              31-MAR-2003; 2003WO-JP004108.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               29-MAR-2002; 2002JP-00132121.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (MOCH ) MOCHIDA PHARM CO LID
                                                                                                                                              Toll-like receptor 4 (Tlr-4)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        209/c
ADF17209 standard; DNA; 25
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (first entry)
                                                                                                                                                                                                                                                                                                 21; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Furusako S, Satoh T,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           WPI; 2003-903121/82.
                                                                                                                                                                                                                                                                       Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WO2003087072-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          12-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             23-OCT-2003.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ADF17209;
                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                 Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 28

ADF17209/
1D ADF17209/
1D ADF17209/
1D ADF17209/
1D ADF17209/
ADF
8888888888
                                                                                                                                                                                                                                                                                                                                                  ઠ
                                                                                                                                                                                                                                                                                                                                                                                         g
```

```
ö
                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           The invention relates to modulating an immune response in an animal. The method of the invention comprises modulating colony stimulating factor-1 (CSF-1) activity in order to modulate the immune response of the animal. Also disclosed is a pharmaceutical composition comprising a modulator of CSF-1 activity and a pharmaceutical carrier. The method or the pharmaceutical correct of the method or the pharmaceutical composition is useful for the prophylactic or therapeutic treatment of bacterially-induced septic shock. The sequences given in records ACC74129-ACC74161 represent primers and probes used in an example from the invention to detect murine genes
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Modulating an immune response in an animal, useful for the prophylactic or therapeutic treatment of bacterially-induced septic shock, by modulating colony stimulating factor-1 (CSF-1) activity in an animal.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Mouse, immunomodulator; antibacterial; immunosuppressive; CSF-1; colony stimulating factor-1; septic shock; TLR; toll-like receptor; interleukin-12; IL-12; HPRT; hypoxanthine phosphoribosyl transferase;
                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                   ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Score 20.2; DB 1; Length 25;
Pred. No. 1e+02;
0; Mismatches 3; Indels
                                         Score 21; DB 1; Length 25; Pred. No. 84;
                                                                                                   0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 25 BP; 6 A; 10 C; 3 G; 6 T; 0 U; 0 Other;
Sequence 25 BP; 12 A; 5 C; 3 G; 5 T; 0 U; 0 Other;
                                 0.6%; Scott No. 100.0%; Pred. No. 0, Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Sester DP;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Probe for detecting murine TLR4 expression.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 TGGTGGCTGTGGAGACAAATCTAGC 580
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   regreereresadacaaarreec 1
                                                                                                                                                           1140 TTTTCTTATATTTCGGATGG 1160
                                                                                                                                                                                                             21 rrrrcrraraarrcccarcc 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Stacey KJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Example; Page 19; 58pp; English.
                                                                                                                                                                                                                                                                                                                                               ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.5%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           03-OCT-2002; 2002WO-AU001348.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             03-OCT-2001; 2001AU-00008071
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAI71235
ID AAI71235 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                         ACC74143/c
ID ACC74143 standard; DNA; 25
                                                                                                                                                                                                                                                                                                                                                                                                                                                         (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  22; Conservative
                                                                                                      Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (UYQU ) UNIV QUEENSLAND.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Hume DA, Sweet MJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2003-381587/36
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Similarity
                                                                             Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO2003028752-A1.
                                                 Query Match
Best Local Simil
Matches 21; C
                                                                                                                                                                                                                                                                                                                                                                                                                                                         11-JUL-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
Best Local S
Matches 22
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                probe; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       556
                                                                                                                                                                                                                                                                                                                                                                                                    ACC74143;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      25
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Mus sp
```

```
The present invention describes an anti-CD14 antibody, which has a function of inhibiting the binding of CD14 to the Toll-like receptor (TLR). The anti-CD14 antibody can specifically recognise the epitope containing the domain from numbers 269-315 in human CD14 of the sequence in AAG68127 or a part of it. Anti-CD14 antibody has antibacterial, immunosuppressive, antipyretic, hypertensive, immunostimulant, haemostatic and vasocropic activities. The antibody together with other polypeptides are applicable in drugs for treating bacterial infection as well as sepsis, fever, hypotension, leukopaenia, thrombopaenia, shock and multi-organ failure. AAG68127 to AAG68137 and AAI71230 to AAI71295
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            represent sequences used in the exemplification of the present invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Anti-CD14 antibody or its fragment inhibiting the binding of CD14 to To-like receptor, applicable in drugs for treating bacterial infection as well as sepsis, fever, hypotension, leukopenia, thrombopenia and shock.
                                                                                                                          Human, Toll like receptor; TLR; CD14; antibody; anti-CD14 antibody; TLK/CD14 binding inhibitor; antibacterial; immunosuppressive; antipyretic; hypertensive; immunostimulant; haemostatic; vasotropic; bacterial infection; sepsis; fever; hypotension; leukopaenia; thrombopaenia; shock; multi-organ failure; ss.
                                                                                         Human Toll like receptor 4 PCR sense primer 2 SEQ ID NO:8.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 4 A; 8 C; 3 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Human TLR4 gene exon 4 amplifying reverse primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      100.0%; Pred. No. 73; ive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Example 2; Page 169; 202pp; Japanese.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            501 CCCATCCAGAGTTTAGCCCT 520
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1 CCCATCCAGAGTTTAGCCCT 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAC84818 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                31-MAR-2000; 2000JP-00099617.
                                                                                                                                                                                                                                                                                                                                                                                                                   2000JP-00356719.
                                                                                                                                                                                                                                                                                                                                                                                                                                   28-MAR-2001; 2001US-00806158.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (MOCH ) MOCHIDA PHARM CO LTD.
                                                                                                                                                                                                                                                                                                                                                           02-APR-2001; 2001WO-JP002869
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (first entry)
                                                    23-JAN-2002 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Furusako S, Mori S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2001-616487/71.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Best Local Similarity
Matches 20, Conser
                                                                                                                                                                                                                                                                                 WO200172993-A1.
                                                                                                                                                                                                                                              Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20-APR-2001
                                                                                                                                                                                                                                                                                                                      04-OCT-2001
                 AAI71235;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AAC84818;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      RESULT 31
ద
```

Shirakawa K, Takahashi T;

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk off, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LDS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS),

Syndrome (SIRS) or acute respiratory distress syndrome (ARDS),

Systematic fibrosis, Sequences AACG476-823 represent PCR primers for and cystic fibrosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                            Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNJ amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; 0; Indels tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 3 A; 4 C; 8 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                   Schwartz DA, Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              2380 AGCACTTCATCCAGAGCCGC 2399
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AGCACTTCATCCAGAGCCGC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        BP
                                                                                                                                                                                                       08-JUN-2000, 2000WO-US015723.
                                                                                                                                                                                                                                              10-JUN-1999; 99US-00329515.
                                                                                                                                                                                                                                                                                    IOWA ) UNIV IOWA RES FOUND.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAC84789 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match 0.5
Best Local Similarity 100.
Matches 20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                         (LORE/) LORENZ E.
                                                                                                                         WO200077204-A1.
                                                                                     Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      20-APR-2001
                                                                                                                                                                 21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             receptor 4.
                                                                                                                                                                                                                                                                                                                                                   Lorenz E,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AAC84789;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 32
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAC84789
      ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Toll
```

ö

Gaps ö

0; Indels

0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73;

WO200077204-A1.

```
methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis.

In pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LDS (lippolysaccharide) induced induced by gram-negative bacteria, LDS (lippolysaccharide) induced inflammatory disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory accounting syndrome (STRS) or acute respiratory distress syndrome (ARDS).

Syndrome (STRS) or acute respiratory distress syndrome (ARDS).
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           TIR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory; disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               The invention relates to human toll receptor 4 (TLR4) nucleic acid and
                                                                                                                                                                                                                                                                                                                                                                  Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0.5%; Score 20; DB 1; Length 20; 00.0%; Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 20 BP; 6 A; 7 C; 2 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  100.08; Pred. M. Hismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   and cystic fibrosis. Sequences AAC84776 amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                        Schwartz DA, Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1612 TCCAGGAAACTTCCTTCCA 1631
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAC84791 standard; DNA; 20 BP
                                                                                                                                             08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                    99US-00329515.
                                                                                                                                                                                                                         (IOWA ) UNIV IOWA RES FOUND (LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
Best Local Similarity 100.
Matches 20, Conservative
                                                                                                                                                                                                                                                                                                                             WPI; 2001-061872/07
                                                            WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
                                                                                                                                                                                    10-JUN-1999;
                       Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20-APR-2001
                                                                                                      21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                         receptor 4.
                                                                                                                                                                                                                                                                                          Lorenz E,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AAC84791;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AAC84791
ID AAC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ઠ
```

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered individuals immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative Sepsis, con pre-term infants, agents which alter TLR4 activity are useful for in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, included by gram-negative bacteria, included induced conditions such as systemmic inflammatory disease conditions such as systemmic inflammatory disease conditions such as systemmic inflammatory disease conditions such as systemmic inflammatory disease, astham arthritis, local and systemic conspiritis, gall bladder disease, preumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                rER4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory; disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                   Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 20 BP; 5 A; 5 C; 5 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          100.0%; Prec. ...
                                                                                                                                                                                                              Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1924 ACCAGAGITICCIGCAATGG 1943
                                                                                                                                                                                                                                                                                                                                                                                        Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ACCAGAGITICCIGCAAIGG 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AAC84798 standard; DNA; 20 BP
                                                                        08-JUN-2000; 2000WO-US015723.
                                                                                                                 99US-00329515.
                                                                                                                                                    UNIV IOWA RES FOUND.
LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       20; Conservative
                                                                                                                                                                                                                Schwartz DA,
                                                                                                                                                                                                                                                      WPI; 2001-061872/07
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
Best Local Similarity
Matches 20; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20-APR-2001
                                                                                                               10-JUN-1999;
                                   21-DEC-2000
                                                                                                                                                                                                                                                                                                                                                         receptor 4.
                                                                                                                                                                                                                   lorenz E,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AAC84798;
                                                                                                                                                       (IOWA)
(LORE/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAC84798
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  셤
```

ö

ö

Gaps

; 0

0; Indels

WO200077204-A1.

21-DEC-2000.

Homo sapiens.

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered inmunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory discress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, leS (liopolysaccharide) induced by gram-negative bacteria, LPS (liopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (STRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                    Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 7 A; 6 C; 5 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                    Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                         Example 1; Page 31; 97pp; English.
                       08-JUN-2000; 2000WO-US015723.
                                                                    99US-00329515
                                                                                                             (IOWA ) UNIV IOWA RES FOUND
                                                                                                                                                                                 Schwartz DA,
                                                                                                                                                                                                                            WPI; 2001-061872/07
                                                                                                                                    (LORE/) LORENZ E.
                                                                    10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                           receptor 4.
                                                                                                                                                                                 Lorenz E,
```

; 0 0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; 0; Indels 100.0%; Pred. No. 73; ative 0; Mismatches 2590 GACGACTCAGAAAAGCCCTG 2609 1 GACGACTCAGAAAGCCCTG 20 Local Similarity 100. les 20; Conservative Query Match Matches ઠ 셤

RESULT 35

Human TLR4 gene exon 4 amplifying reverse primer.

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss. 

99US-00329515 10-JUN-1999;

(IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E.

Schwartz DA, Schutte BC; Lorenz E,

WPI; 2001-061872/07

Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll receptor 4.

Example 1; Page 31; 97pp; English.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LDS (lipopolygaccharide) induced chronic airway disease conditions such as systematic inflammatory response inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), systematic inflammatory disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC64776-823 represent PCR primers for and cystic fibrosis. Sequences AAC84776. amplifying the exons of human TLR4 gene 

Sequence 20 BP; 2 A; 5 C; 6 G; 7 T; 0 U; 0 Other;

Gaps ö 0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; 0; Indels 100.0%; Prea. ... Matches 20; Conservative Local Similarity Ouery Match

ö

2590 GACGACTCAGAAAGCCCTG 2609 20 GACGACTCAGAAAAGCCCTG 1 g ઠે

ö

Gaps

AAC84775 standard; DNA; 20 BP AAC84775; RESULT 36 AAC84775/ 

Human TLR4 gene amplifying primer 2R. (first entry) 20-APR-2001

TLR4, toll receptor 4, innate immunity, gram-negative bacteria; sepsis, respiratory distress syndrome; LPS, lipopolysaccharide, asthma, ARDS, chronic airway disease; arthritis; inflammatory disease; SIRS; human; Bystematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

Ното варіепв

WO200077204-A1.

21-DEC-2000.

08-JUN-2000; 2000WO-US015723

99US-00329515. 10-JUN-1999;

(IOWA ) UNIV IOWA RES FOUND.

10001863-3.81.rng

```
(IOWA ) UNIV IOWA RES FOUND.
(LORE/) LORENZ E.
          WPI; 2001-061872/07
                                                                                                                                                              WO200077204-A1
                                                                                                                                                         Homo sapiens.
                                                                                                                                                                              10-JUN-1999;
                                                                                                                                                                   21-DEC-2000.
                                                                                                                                                                                           Lorenz E,
     Lorenz E,
                                                                                                                       AAC84785;
                       receptor
                                                                                      Query Match
Best Local 8
                                                                                           tches
                                                                                                             RESULT 37
                                                                                                               쉱
```

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the comparint of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, infanta, sepsis conducted by gram-negative bacteria, info control airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), constructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic alrway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                 Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNJ amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 73;
tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Sequence 20 BP; 5 A; 5 C; 4 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1190 TGGACAGTTTCCCACATTGA 1209
                                                                                                                                                                                                           Example 1; Page 31; 97pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1 TGGACAGTTTCCCACATTGA 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      08-JUN-2000; 2000WO-US015723
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (IOWA ) UNIV IOWA RES FOUND
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AAC84792 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Lorenz E, Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Conservative
                      WPI; 2001-061872/07
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Best Local Similarity
Matches 20; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WO200077204-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AAC84792;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 LORE/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 38
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAC84792
ò
                                                                                                                                                                                                                                                                                                                                              The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 uncleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or amaliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), prolomophritis, gall bladder disease, pneumonia, bronchitis, chronic constructive pulmonary disease, local gram-negative bacterial infection amplifying the human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                           Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Sequence 20 BP; 2 A; 5 C; 6 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              2590 GACGACTCAGAAAAGCCCTG 2609
                                                                                                                                                                                                                                                                                                      Example 1; Page 29; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAC84785 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 20-APR-2001 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              20; Conservative
                                                           Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Local Similarity
  (LORE/) LORENZ E.
```

WPI; 2001-061872/07

Schwartz DA, Schutte BC;

ö

Gaps

methods to identify polymorphisms at the human TiR4 locus and to identify indiduals at risk of, or having, an indication associated with altered innate immunity. A variant TiR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TiR4 gene. Since the presence of TiR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TiR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, incl. activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response inflammatory disease conditions such as systematic inflammatory response systemome (SRRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic conditions to present port prevential infection and systic fibrosis. Sequences AAC84776-823 represent PCR primers for invention relates to human toll receptor 4 (TLR4) nucleic acid and altered innate immunity involves detecting or determining whether DN amplified from a biological sample encodes a portion of variant toll Identifying humans at risk of, or having indication associated with amplifying the exons of human TLR4 gene Example 1; Page 31; 97pp; English. receptor 

Sequence 20 BP; 3 A; 3 C; 6 G; 8 T; 0 U; 0 Other;

; 0 0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; Live 0; Mismatches 0; Indels 20; Conservative Best Local Similarity Query Match **fatches** 

2017 TGCCTGTGCTGAGTTTGAAT 2036 TGCCTGTGCTGAGTTTGAAT 20 g 8

AAC84799 standard; DNA; 20 BP AAC84799; RESULT 39 AAC84799 

20-APR-2001 (first entry)

Human TLR4 gene exon 4 amplifying forward primer.

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pheumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

Homo sapiens

WO200077204-A1.

21-DEC-2000.

08-JUN-2000; 2000WO-US015723.

10-JUN-1999;

(IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E.

Schutte BC; Lorenz E, Schwartz DA,

WPI; 2001-061872/07.

Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll

```
methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis chronic airway disease, astham, arthritis, local and systemic inflammatory disease conditions such as systematic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AACH4776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                     The invention relates to human toll receptor 4 (TLR4) nucleic acid and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 20 BP; 8 A; 4 C; 5 G; 3 T; 0 U; 0 Other;
                                                          Example 1; Page 31; 97pp; English.
```

Score 20; DB 1; Length 20; Pred. No. 73; 0; Indels 0.5%; bcc. 100.0%; Pred. No. ... 2658 AATTGGCAGGAAGCAACATC 2677 20; Conservative Similarity Query Match Local Matches 셤 à

1 AATTGGCAGGAAGCAACATC 20

ö

Gaps

ö

Gaps

ö

AAC84825 standard; DNA; 20 (first entry) 20-APR-2001 AAC84825; RESULT 40 AAC84825/ 

BP

Reverse primer derived form human TLR4 gene exon 4.

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress, syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic alrway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

Homo sapiens

WO200077204-A1.

21-DEC-2000.

08-JUN-2000; 2000WO-US015723.

(IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E.

.0-JUN-1999; 99US-00329515

Lorenz E, Schwartz DA, Schutte BC;

WPI; 2001-061872/07

Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll receptor 4.

Example 1; Page 33; 97pp; English.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 unclaic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation lis associated with gram-negative sepsis. Coverity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, proceed by gram-negative bacteria, sepsis induced a riway disease, asthma, arthritis, local and systemic chronic airway disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchtis, chronic and cystic fibrosis. The present sequence represents a reverse primer and cystic fibrosis. The present sequence represents a reverse primer constructive dorm exon 4 of the human TLR4 gene, used in multi-tissue cDNA expression screen 

Seguence 20 BP; 4 A; 6 C; 4 G; 6 T; 0 U; 0 Other;

DB 1; Length 20; 0; Indels 0.5%; Score 20; DB 100.0%; Pred. No. 73; ive 0; Mismatches 448 ATGGGCATATCAGAGCCTA 467 20 ATGGGGCATATCAGAGCCTA 1 Local Similarity 100. ses 20; Conservative Query Match Matches à 셤

RESULT 41

AAC84812 standard; DNA; 20 AAC84812 

H

AAC84812;

20-APR-2001 (first entry)

Human TLR4 gene exon 4 amplifying reverse primer.

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic alrway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

Homo sapiens.

WO200077204-A1.

21-DEC-2000.

08-JUN-2000; 2000WO-US015723

99US-00329515 10-JUN-1999;

(IOWA ) UNIV IOWA RES FOUND (LORE/) LORENZ E.

Schutte BC; Schwartz DA, ы Lorenz

WPI; 2001-061872/07.

altered innate immunity involves detecting or determining whether DN amplified from a biological sample encodes a portion of variant toll Identifying humans at risk of, or having indication associated with receptor 4.

Example 1; Page 31; 97pp; English.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and

methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory disease, patent as arthritis, local and systemic syndrome (SIRS) or acute respiratory distress syndrome (ARDS), concerning and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene ö Gaps ö 0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; ive 0; Mismatches 0; Indels Sequence 20 BP; 7 A; 3 C; 8 G; 2 T; 0 U; 0 Other; 20; Conservative Similarity Query Match Local Matches \$

1656 TIGACCTICCIGGACCTCIC 1675 20 TTGACCTTCCTGGACCTCTC

ò

ö

Gaps

; 0

AAC84774 standard; DNA; 20 BP RESULT 42 AAC84774 

AAC84774;

(first entry) 20-APR-2001

Human TLR4 gene amplifying primer 2F.

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

Homo sapiens.

WO200077204-A1

21-DEC-2000.

08-JUN-2000; 2000WO-US015723.

10-JUN-1999;

(IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E. Schwartz DA, Schutte BC; Lorenz E,

WPI; 2001-061872/07

Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNI amplified from a biological sample encodes a portion of variant toll

Example 1; Page 29; 97pp; English.

receptor

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered inmane immunity. A variant TLR4 nucleic acid is useful as a diagnostic

```
reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, per-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory all bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84772-775 represent PCR primers for amplifying the human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 20 BP; 4 A; 4 C; 6 G; 6 T; 0 U; 0 Other;
         88888888888888888888888888
```

```
Score 20; DB 1; Length 20;
Pred. No. 73;
                        0; Indels
                        0; Mismatches
                                               2467 TCATTGTCCTGCAGAAGGTG 2486
         100.08;
  0.5%;
                        20; Conservative
            Similarity
Query Match
              Local
                       Matches
                                                                    g
```

ö

Gaps .. 0

1 rcarrercrecadadere 20

```
AAC84804 standard; DNA; 20 BP
   AAC84804;
RESULT 43
```

20-APR-2001 (first entry)

Human TLR4 gene exon 4 amplifying reverse primer.

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

Homo sapiens

WO200077204-A1.

21-DEC-2000.

08-JUN-2000; 2000WO-US015723.

10-JUN-1999; 99US-00329515

(IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E.

Schutte Lorenz E, Schwartz DA,

BG

WPI; 2001-061872/07.

Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll receptor 4.

Example 1; Page 31; 97pp; English.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome

```
ö
in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), performed pythitis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC847/6-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                      Gaps
                                                                                                                                                                       ö
                                                                                                                                                Score 20; DB 1; Length 20;
Pred. No. 73;
                                                                                                                                                                      0; Indels
                                                                                                                           Sequence 20 BP; 7 A; 5 C; 4 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                          Human TLR4 gene exon 4 amplifying reverse primer.
                                                                                                                                          0.5%; Scc.
100.0%; Pred. No. ...
                                                                                                                                                                                           733 TTTATTGCACAGACTTGCGG 752
                                                                                                                                                                                                                  20 TTTATTGCACAGACTTGCGG 1
                                                                                                                                                                                                                                                                         AAC84813 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                   20-APR-2001 (first entry)
                                                                                                                                           Query Match
Best Local Similarity 100."
Matches 20; Conservative
                                                                                                                                                                                                                                                                                               AAC84813;
                                                                                                                                                                                                                                                     RESULT 44
                                                                                                                                                                                                                                                                 AAC84813/
   888888888888888888
                                                                                                                                                                                                                                                                                      ò
                                                                                                                                                                                                                셤
```

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic.alrway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

Homo sapiens

WO200077204-A1.

21-DEC-2000.

08-JUN-2000; 2000WO-US015723.

10-JUN-1999; 99US-00329515.

(IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E.

Lorenz E, Schwartz DA, Schutte BC;

WPI; 2001-061872/07.

Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll receptor 4.

Example 1; Page 31; 97pp; English.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, lepsis

pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TER4 gene

Sequence 20 BP; 4 A; 5 C; 4 G; 7 T; 0 U; 0 Other;

ö

Gaps

ô

DB 1; Length 20; 73; 0; Indels

0.5%; Scc... 100.0%; Pred. No. .c.

20, Conservative

Similarity

Query Match Best Local ( Matches

8×333333

2659 ATTGGCAGGAAGCAACATCT 2678

ð 셤

ö

20 ATTGGCAGGAAGCAACATCT 1

```
inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, preumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                        Gaps
                                                                                                                                                                        .;
0
chronic airway disease, asthma, arthritis, local and systemic
                                                                                                                                          DB 1; Length 20; 73;
                                                                                                                                                                        0; Indels
                                                                                                             Sequence 20 BP; 6 A; 4 C; 7 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                       Human TLR4 gene exon 4 amplifying reverse primer.
                                                                                                                                                         100.0%; Pred. No. 73; ive 0; Mismatches
                                                                                                                                           Score 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Schutte BC;
                                                                                                                                                                                                    1788 IGTCTGAACTCCCTCCAGGT 1807
                                                                                                                                                                                                                     20 TGTCTGAACTCCCTCCAGGT 1
                                                                                                                                                                                                                                                                                                      BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      99US-00329515
                                                                                                                                           0.5%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 UNIV IOWA RES FOUND LORENZ E.
                                                                                                                                                                                                                                                                                                      AAC84820 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                             (first entry)
                                                                                                                                                                        20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Lorenz E, Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2001-061872/07
                                                                                                                                                         Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (IOWA ) UNIV
                                                                                                                                                                                                                                                                                                                                                               20-APR-2001
                                                                                                                                                                                                                                                                                                                                  AAC84820;
                                                                                                                                              Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (LORE/)
                                                                                                                                                                         Matches
                                                                                                                                                                                                                                                                           888888888
                                                                                                                                                                                                      à
                                                                                                                                                                                                                               임
```

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered individuals at risk of separatory of electing a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LDS (lipopolysaccharide) induced chronic airway disease conditions such as systematic inflammatory response inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), syndrome (SIRS) at all bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for
                                                                                                                                                                               TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic alrway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DN amplified from a biological sample encodes a portion of variant toll
                                                                                                                                          Human TLR4 gene exon 4 amplifying reverse primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Schutte
                   ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                           08-JUN-2000; 2000WO-US015723
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              IOWA ) UNIV IOWA RES FOUND.
                 AAC84822 standard; DNA; 20
                                                                                                  (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                            WO200077204-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                    Homo sapiens.
                                                                                                  20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                     21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Lorenz E,
                                                             AAC84822;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              receptor
AAC84822
```

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, per-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lippoblysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic chronic airway disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS),

Example 1; Page 31; 97pp; English.

10001863-3.sl.rng

ö

Gaps

ö

```
methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered innainty. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, arthritis, local and systemic chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephitis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84772-775 represent PCR primers for amplifying the human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               The invention relates to human toll receptor 4 (TLR4) nucleic acid and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                   DB 1; Length 20; 73;
                                                                                                                                                                                                                             Indels
                                                                         Sequence 20 BP; 6 A; 4 C; 6 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                      100.0%; Pred. ...
amplifying the exons of human TLR4 gene
                                                                                                                                                   Score 20;
Pred. No.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Human TLR4 gene amplifying primer 1R.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Schutte BC;
                                                                                                                                                                                                                                                                                                   2725 CTGGGTCCAACACTTGTTCA 2744
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Example 1; Page 29; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                 20 CTGGGTCCAACACTTGTTCA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAC84773 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           08-JUN-2000; 2000WO-US015723.
                                                                                                                                                   0.5%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (IOWA ) UNIV IOWA RES FOUND.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20-APR-2001 (first entry)
                                                                                                                                                                                  Best Local Similarity 100.
Matches 20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Lorenz E, Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WO200077204-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Homo sapiens,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 21-DEC-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               receptor 4.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAC84773;
                                                                                                                                                       Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AAC84

                                                                                                                                                                                                                                                                                                      8
```

```
methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 mucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory discress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LDS (lipopolysaccharide) induced contains arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory disease, asthma, arthritis, local and systemic syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AACS4776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                    TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; preumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; se.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        The invention relates to human toll receptor 4 (TLR4) nucleic acid and
                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    altered innate immunity involves detecting or determining whether DN amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           identifying humans at risk of, or having indication associated with
                                                 ö
          Length 20;
                                                 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 20 BP; 5 A; 7 C; 3 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                 Human TLR4 gene exon 4 amplifying forward primer.
          Score 20; DB 1;
Pred. No. 73;
0.5%; bcc.
100.0%; Pred. No. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Lorenz E, Schwartz DA, Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Example 1; Page 31; 97pp; English.
                                                                                        909
                                                                                                              20 AGAGAACTICCCCATIGGAC 1
                                                                                        587 AGAGAACTICCCCATIGGAC
                                                                                                                                                                                                                               ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     .0-JUN-1999; 99US-00329515.
                                                                                                                                                                                                                           AAC84781 standard; DNA; 20
                                                                                                                                                                                                                                                                                                        (first entry)
                            Local Similarity 100.
nes 20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2001-061872/07
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens
                                                                                                                                                                                                                                                                                                        20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            receptor 4.
                                                                                                                                                                                                                                                                  AAC84781;
            Query Match
                                                 Matches
                                                                                                                                                                                      RESULT 48
                                                                                                                                                                                                      AAC84781
                                                                                                                                                                                                                                                                  ઠ
```

DB 1; Length 20; 73;

0.5%; Score 20; 100.0%; Pred. No.

Query Match Best Local Similarity

Seguence 20 BP; 4 A; 4 C; 6 G; 6 T; 0 U; 0 Other;

1 AAGCCGAAAGGTGATTGTTG 20

셤

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innunte immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis.

Dieserty of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (SIRS) or acute respiratory distress syndrome (ABRS), choonic obstructive pulmonary disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, agam-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
       Gaps
       ö
       Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Seguence 20 BP; 6 A; 2 C; 7 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Human TLR4 gene exon 4 amplifying forward primer.
       ö
           Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Example 1; Page 31; 97pp; English.
                                                                             700 ACTIGGACCTTTCCAGCAAC 719
                                                                                                                                               20
           ô
                                                                                                                                                   1 ACTIGGACCTITCCAGCAAC
                                                                                                                                                                                                                                                                                                                         AAC84795 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               99US-00329515
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (IOWA ) UNIV IOWA RES FOUND.
(LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (first entry)
           Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2001-061872/07
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO200077204-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   21-DEC-2000.
           20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Lorenz E,
                                                                                                                                                                                                                                                                                                                                                                                                   AAC84795;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            receptor
              Matches
                                                                                                                                                                                                                                                          RESULT 45
AAC8 4795
AAC8 4
                                                                                                                                                   셤
                                                                                 ð
```

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered individuals at tisk of, or having, an indication as a diamostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LBS (lipopolysaccharide) induced chonic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (STRS) or acute respiratory distress syndrome (ATRS), or acute respiratory distress syndrome (ATRS), obstructive pulmonary disease, neumonia, bronchitis, considered form expensed formar aforward primer and cystic fibrosis. The present sequence represents a forward primer and cystic fibrosis. The present sequence represents a forward primer and cystic fibrosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                            TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; tive 0; Mismatches 0; Indels
                                                                                                                                                        Forward primer derived form human TLR4 gene exon 1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 20 BP; 7 A; 4 C; 7 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Lorenz E, Schwartz DA, Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Example 1; Page 33; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 15 GCTCACAGAAGCAGTGAGGA 34
                                   BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         .0-JUN-1999; 99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (IOWA ) UNIV IOWA RES FOUND.
(LORE/) LORENZ E.
                                     20
                                                                                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            20; Conservative
                                   AAC84824 standard; DNA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2001-061872/07
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    expression screen
                                                                                                                                                                                                                                                                                                                                                                                                 WO200077204-A1
                                                                                                                                                                                                                                                                                                                                                         Homo sapiens
                                                                                                                       20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                         21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      receptor 4.
                                                                               AAC84824;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Matches
                   AAC8482
RESULT
```

20; Conservative

Similarity

Query Match Best Local &

Matches

1 gcrcacagaagcagrgagga 20

g 8

ö

Gaps

ö

0; Indels

Score 20; DB 1; Length 20; Pred. No. 73;

toll receptor 4; innate immunity; gram-negative bacteria; sepsis;

Human TLR4 gene exon 4 amplifying reverse primer.

(first entry)

20-APR-2001

AAC84815 standard; DNA; 20 BP.

AAC84815/c

respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; STRS; human; systematic inflammatory response syndrome; pyclonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LBS (lipopolysaccharide) induced chronic airway disease conditions authritis, local and systemic inflammatory disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory disease, preumonia, bronchitis, ornonic obstructive pulmonary disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, pneumonia, bronchitis, chronic and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                     TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                               Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 20 BP; 5 A; 3 C; 8 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Example 1; Page 31; 97pp; English.
                                         AAC84780 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                99US-00329515
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (IOWA ) UNIV IOWA RES FOUND
                                                                                                                          (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Lorenz E, Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                   WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              10-JUN-1999;
                                                                                                                          20-APR-2001
                                                                                                                                                                                                                                                                                                                                                              Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                               21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         receptor 4.
                                                                                   AAC84780;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
RESULT 51
                     AAC84780
```

Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll

Example 1; Page 31; 97pp; English.

receptor 4.

corenz E, Schwartz DA, Schutte BC;

WPI; 2001-061872/07.

99US-00329515.

10-JUN-1999;

(IOWA ) UNIV IOWA RES FOUND.

(LORE/) LORENZ E.

38-JUN-2000; 2000WO-US015723.

WO200077204-A1. Homo sapiens.

11-DEC-2000.

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered intate immunity. A variant TLR4 is acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepplis. Correspondently of sepals, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepplis induced by gram-negative bacteria, sepplis induced chronic airway disease, asthma, arthritis, local and systemic chronic airway disease, conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS). Pyelonephitits, gall bladder disease, pneumonia, bronchitis, chronic cobstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences Ascass gram-negative bacterial infection and cystic fibrosis. Sequences Ascass gram-negative bacterial infection and cystic fibrosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; ive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 20 BP; 7 A; 6 C; 5 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            2074 CGGTCCTCAGTGTGTTGTA 2093
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            20:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Best Loc
Matches
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      0.5%; Score 20; DB 1; Length 20; 00.0%; Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0; Indels
```

```
Gaps
ö
                                     20 cecrcicaererectiona
                                                                                         AAC84816 standard; DNA; 20 BP
                                                                             AAC84816/c
ID AAC84
XX
                                                                     RESULT 53
                                     g
```

ö

RESULT 52

0; Mismatches

100.0%;

20; Conservative

Best Local Similarity

Matches

ò

557 GGTGGCTGTGGAGACAAATC 576

GGTGGCTGTGGAGACAAATC

-

20

```
CTTGCTGGCTGCATAAA 2161
                                                                                                                                                                                                                                         20 chrchigerechicaraaa 1
                                                                                                                                                                                                                                                            AAC84819 standard; DNA; 20
                                                                                                                                                                                                                  Query Match
Best Local Similarity 100.'
Marches 20, Conservative
                                                                                                        WPI; 2001-061872/07
                                                       WO200077204-A1
                                                                              10-JUN-1999;
                                                 Homo sapiens,
                                                                                                                                                                                                                                  2142 CTT
                                                               21-DEC-2000.
                                                                                                                           receptor 4.
                                                                                                Lorenz E,
                                                                                                                                                                                                                                                    RESULT 54
                                                                                                                                                                                                                                                        AAC84819/
ID AAC8
XX
AC AAC8
XX
DT 20-A
ò
```

```
TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic alrway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                  Human TLR4 gene exon 4 amplifying reverse primer.
                                                                                                                                                                                                                                                                                                                                                                                                     08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (IOWA ) UNIV IOWA RES FOUND.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                  WO200077204-A1.
                                                                                                                                                                                                                                                       Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                          10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                       21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Lorenz E,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          receptor
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (LORE/)
ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 uncleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLRA mutation is associated with gram-negative sepsis, creatity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AG84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                               TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DN amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 73;
ive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Seguence 20 BP; 7 A; 4 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                     Human TLR4 gene exon 4 amplifying reverse primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (IOWA ) UNIV IOWA RES FOUND (LORE/) LORENZ E.
                                                  20-APR-2001 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Schwartz DA,
AAC84816;
```

Schwartz DA, Schutte BC;

LORENZ E.

99US-00329515.

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for in pre-term infants, agents which alter TLR4 activity are useful for creating or ameliocrating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LBS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic chronic airway disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic and cystic fibrosis. Sequences AACSA represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ő
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Sequence 20 BP; 3 A; 7 C; 5 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  100.0%; Prec. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2484 GTGGAGAGACCCTGCTCAG 2503
Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            GTGGAGAAGACCCTGCTCAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 멂
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AAC84794 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 55
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAC84794
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 셤
```

20-APR-2001 (first entry)

AAC84819;

```
TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                           Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNJ amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                         Example 1; Page 31; 97pp; English.
                                                                                                                                                           08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                         IOWA ) UNIV IOWA RES FOUND
                                                                                                                                                                                                                                               Lorenz B, Schwartz DA,
                                                                                                                                                                                                                                                                     WPI; 2001-061872/07
                                                                                                                                                                                                                     LORE/) LORENZ E.
                                                                                                            WO200077204-A1
                                                                                                                                                                                 10-JUN-1999;
                                                                                    Homo sapiens.
                                                                                                                                    21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                   receptor 4.
```

Schutte BC;

99US-00329515.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 uncleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for in preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, sepsis induced by gram-negative bacteria, lDS (lipopolysaccharide) induced chronic airway disease asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephitis, gall bladder disease, pneumonia, bronchtis, chronic and cystic fibroais. Sequences ARG4776-823 represent PCR primers for amplifying the exons of human TLR4 gene Gaps ö 0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; 0; Indels Sequence 20 BP; 6 A; 3 C; 8 G; 3 T; 0 U; 0 Other; 100.0%; Preα. ... ive 0; Mismatches Conservative Local Similarity les 20; Conserv Query Match Best Loc Matches

2210 CCAGGATGAGGACTGGGTAA 2229 CCAGGATGAGGACTGGGTAA 20

셤 δ

AAC84797 standard; DNA; 20 BP AAC84797; RESULT 56 

(first entry) 20-APR-2001

Human TLR4 gene exon 4 amplifying forward primer.

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human;

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, conserved to TLR4 mutation is associated with gram-negative sepsis, conserved by gram-negative bacteria, sepsis in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis conducted by gram-negative bacteria, sepsis conducted by gram-negative bacterial, LDS (lippoblysaccharide) induced chronic airway disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pelonophritis, gall bladder disease, pneumonia, bronchisis, chronic ond cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene mmatory response syndrome; pyelonephritis; bronchitis; y distress syndrome; gall bladder disease; pneumonia; antibacterial; antiinflammatory; PCR primer; ss. Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll Sequence 20 BP; 4 A; 4 C; 6 G; 6 T; 0 U; 0 Other; Schwartz DA, Schutte BC; Example 1; Page 31; 97pp; English. 08-JUN-2000; 2000WO-US015723. 10-JUN-1999; 99US-00329515. (IOWA ) UNIV IOWA RES FOUND WPI; 2001-061872/07. acute respiratory cystic fibrosis; a LORENZ E. WO200077204-A1 21-DEC-2000. receptor 4. Lorenz E, (LORE/) 

Gaps ö 0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; 0; Indels 100.0%; Pred. No. 73; ive 0; Mismatches 2467 TCATTGTCCTGCAGAAGGTG 2486 1 TCATTGTCCTGCAGAAGGTG Query Match 0.5 Best Local Similarity 100. Matches 20; Conservative ઠે 셤

ö

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss. Human TLR4 gene exon 4 amplifying reverse primer. 

(first entry)

20-APR-2001

AAC84809;

AAC84809 standard, DNA; 20 BP

RESULT 57 AAC84809

99US-00329515.

WO200077204-A1

10001863-3.sl.rng

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, seventity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory disease. Sequences pneumonia, bronchitis, chronic pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                       Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 20 BP; 3 A; 5 C; 5 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                             Lorenz E, Schwartz DA, Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Example 1; Page 31; 97pp; English.
                                                                                                                                                08-JUN-2000; 2000WO-US015723
                                                                                                                                                                                                                             (IOWA ) UNIV IOWA RES FOUND (LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                      WPI; 2001-061872/07
                                                           WO200077204-A1.
                                                                                                                                                                                        .0-JUN-1999;
                     Homo sapiens
                                                                                                    21-DEC-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                receptor 4.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AAC84811;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Matches
8
```

```
Gaps
                         ;
0
Score 20; DB 1; Length 20;
Pred. No. 73;
                        0; Indels
       100.0%; Prea. ...
                                                 1351 TTGGGACAACCAGCCTAAAG 1370
                                                                        20 rreceachaccaccraade 1
  0.5%;
                          20; Conservative
              Local Similarity
```

ö

Human TLR4 gene exon 4 amplifying reverse primer. AAC84811 standard; DNA; 20 BP. (first entry) 20-APR-2001 AAC84811/ IID AAC8 XX AC XX AC XX XX XX XX XX XX XX XX TLR4 XW TLR4 XW CHTO KW CHTO KW CHTO KW CHTO KW CHTO KW CHTO KW COST XX COST

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.

WO200077204-A1

21-DEC-2000.

Homo sapiens.

Homo sapiens

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered inmunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis. Severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LDS (lippol)saccharide) induced chronic airway disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), cypelonephritis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                             Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; cive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 20 BP; 6 A; 6 C; 5 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               amplifying the exons of human TLR4 gene
                                                                                                                                                                                                       Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1568 IGGCTIGICCAGICICGAAG 1587
                                                                                                                                                                                                                                                                                                                                                                                Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TGGCTTGTCCAGTCTCGAAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AAC84788 standard; DNA; 20 BP
                                                                       08-JUN-2000; 2000WO-US015723.
                                                                                                           99US-00329515.
                                                                                                                                                 UNIV IOWA RES FOUND
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20-APR-2001 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1 Similarity 100.
20; Conservative
                                                                                                                                                                                                         Schwartz DA,
                                                                                                                                                                                                                                             WPI; 2001-061872/07.
                                                                                                                                                                     LORENZ E.
                                                                                                             10-JUN-1999;
                                 21-DEC-2000
                                                                                                                                                                                                                                                                                                                                              receptor 4.
                                                                                                                                                                                                           Lorenz E,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAC84788;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match
Best Local S
                                                                                                                                                 IOWA )
                                                                                                                                                                     LORE/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Best Loc
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 59
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AAC84788
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       엄
```

99US-00329515

```
08-JUN-2000; 2000WO-US015723.
          Lorenz E,
                                                                     AAC84793;
                                                   Query Match
                                                      Matches
                                                               RESULT 60
                                                                 AAC84793
ઠે
                                                           g
```

```
10-JUN-1999;
     8X4CCCCCCCCCCX8XFFFFFXXXXFFFFXXXXXF8
                                                                                                                                                                                                                                                                                                                                                                                                                                                              The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered inmunity. A variant TLR4 mucleic acid is useful as a diagnostic for individuals associated with gram-negative sepsis, pre-term delivery and respiratory distress syndrome presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions auch as systematic inflammatory response syndrome (STRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                          Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 20 BP; 5 A; 4 C; 5 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                  Schwartz DA, Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                         Example 1; Page 31; 97pp; English.
                         08-JUN-2000; 2000WO-US015723.
                                                                        99US-00329515
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0.5%;
                                                                                                                       (IOWA ) UNIV IOWA RES FOUND (LORE/) LORENZ E.
                                                                                                                                                                                                                                                 WPI; 2001-061872/07
                                                                      10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                           receptor 4.
```

```
Gaps
                              ő
Score 20; DB 1; Length 20;
Pred. No. 73;
                             0; Indels
             100.0%; Pred. No. 73; vative 0; Mismatches
                                                            1541 CAGAGTTGCTTTCAATGGCA 1560
                                                                        1 CAGAGTTGCTTTCAATGGCA 20
                              20; Conservative
                 Local Similarity
```

ö

TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss. Human TLR4 gene exon 4 amplifying forward primer. AAC84793 standard; DNA; 20 BP 20-APR-2001 (first entry)

WO200077204-A1 Homo sapiens 21-DEC-2000.

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchiis, chronic constructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AACH4776-823 represent PCR primers for amplifying the exons of human TLR4 gene TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss. Gaps Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNJ amplified from a biological sample encodes a portion of variant toll ö 0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; 0; Indels Sequence 20 BP; 2 A; 5 C; 6 G; 7 T; 0 U; 0 Other; Human TLR4 gene exon 4 amplifying reverse primer. 100.0%; Pred. No. Lorenz E, Schwartz DA, Schutte BC; 2074 CGGTCCTCAGTGTGCTTGTA 2093 Example 1; Page 31; 97pp; English. 1 CGGTCCTCAGTGTGCTTGTA 20 AAC84817 standard; DNA; 20 BP 08-JUN-2000; 2000WO-US015723 99US-00329515. (IOWA ) UNIV IOWA RES FOUND. (LORE/) LORENZ E. (IOWA ) UNIV IOWA RES FOUND. (first entry) 20; Conservative WPI; 2001-061872/07. Similarity WO200077204-A1. Homo sapiens. 10-JUN-1999; 20-APR-2001 21-DEC-2000. receptor 4. AAC84817; Query Match Local Matches RESULT 61 AAC84817/ g ò

```
10-JUN-1999;
                                                                20-APR-2001
                                                                               Homo sapiens
                                                                                    21-DEC-2000.
            receptor 4.
                                                             AAC84814;
                                            Query Match
                                              Matches
                                                         ઠે
                                                    g
```

Matches RESULT 63 AAC8477 ò The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication associated with altered individuals at risk of, or having, an indication acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, con pre-term infants, agents which alter TLR4 activity are useful for in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, infa (thoused conditions such as systematic inflammatory disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory is gandrome (SIRS) or acute respiratory distress syndrome (ARDS), conspirative pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-821 represent PCR primers for amplifying the exons of human TLR4 gene ö TLR4, toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss. Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll Gaps ; DB 1; Length 20; 73; 0; Indels Sequence 20 BP; 4 A; 7 C; 4 G; 5 T; 0 U; 0 Other; Human TLR4 gene exon 4 amplifying reverse primer. 0.5%; Scor. 100.0%; Pred. No. ... Lorenz E, Schwartz DA, Schutte BC; Schutte BC 2250 TTAGAAGAAGGGGTGCCTCC 2269 Example 1; Page 31; 97pp; English. 20 TTAGAAGAAGGGGTGCCTCC 1 AAC84814 standard; DNA; 20 BP 08-JUN-2000; 2000WO-US015723. 99US-00329515, IOWA RES FOUND. (first entry) 20; Conservative Lorenz E, Schwartz DA, Local Similarity WPI; 2001-061872/07 (IOWA ) UNIV IOWA (LORE/) LORENZ E. (LORE/) LORENZ E. WO200077204-A1.

```
The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered inmunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, covertive feapsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, local and systemic circlammatory disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory response syndrome (STRS) or acute respiratory distress syndrome (RTRS).

Syndrome (STRS) or acute respiratory distress syndrome (RTRS), chronic obstructive pulmonary disease, including a present to the proposition obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AACG8776-823 represent PCR primers for
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; esystematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                           Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNA amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 20 BP; 3 A; 6 C; 5 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    and cystic fibrosis. Sequences AAC84776 amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Human TLR4 gene amplifying primer 1F.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Schutte BC;
                                                                                                                                                                                                                                                             Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1947 AAGGACCAGAGGCAGCTCTT 19
20 AAGGACCAGAGGCAGCTCTT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            08-JUN-2000; 2000WO-US015723
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (IOWA ) UNIV IOWA RES FOUND
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AAC84772 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Lorenz E, Schwartz DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2001-061872/07.
WPI; 2001-061872/07
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WO200077204-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           10-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AAC84772;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                      receptor
```

```
altered innate immunity involves detecting or determining whether DN
amplified from a biological sample encodes a portion of variant toll
Identifying humans at risk of, or having indication associated with
                                       Example 1; Page 29; 97pp; English.
                        receptor
```

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 uncleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, sepsis induced by gram-negative bacteria, sepsis infection by gram-negative bacteria, sepsis inflammatory disease, asthma, arthritis, local and systemic chronic airway disease, conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchifis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84772-775 represent PCR primers for amplifying the human TLR4 gene

Sequence 20 BP; 6 A; 4 C; 6 G; 4 T; 0 U; 0 Other;

0.5%;

Query Match

Score 20; DB 1; Length 20; Pred. No. 73; 0; Indels 0; Mismatches 448 ATGGGGCATATCAGAGCCTA 467 100.08; 20; Conservative Local Similarity Matches ઠે

ö

Gaps

ö

1 ATGGGGCATATCAGAGCCTA 20 셤

AAL41015 standard; DNA; 20 AAL41015; RESULT 64 AAL41015 

뭠

11-OCT-2002 (first entry)

Anti-CD14 monoclonal antibody related oligonucleotide #3.

Immunosuppressive; antibacterial; anti-CD14 antibody; epitope; sepsis; human CD14; ds.

Unidentified.

WO200242333-A1

30-MAY-2002.

28-SEP-2001; 2001WO-JP008563

22-NOV-2000; 2000JP-00356719

(MOCH ) MOCHIDA PHARM CO LTD

Mori Furusako S, Shirakawa K,

ŝ

WPI; 2002-454920/48.

Anti-CD14 monoclonal antibody which inhibits CD14/T lymphocyte receptor binding by specifically recognizing epitope in human CD14 domain to prevent interaction and suppress cell activation, useful for treating

Example 2; Page 46; 156pp; Japanese

The invention relates to an anti-CD14 antibody which can specifically

ö disease through the administration of a Toll-like receptor-4 (TLR-4) inhibitor to a mammal. The TLR-4 protein has been linked to several disease such as atherosclerosis, restenosis, inflammation and other vascular diseases. Compounds which inhibit the activity of TLR-4, through the inhibition of its receptor, may have vasotropic, antiatrosclerotic, thrombolyvic, cardiant and antiinflammatory activities. This may also be achieved through antisense therapy or gene therapy. The method or the system of the invention may therefore be useful for inhibiting or treating a vascular disease, for example atherosclerosis, transplant atherosclerosis, vein-graft atherosclerosis, thrombosis, stent restenosis, angloplasty restenosis, or inflammation and other heart disease. The present sequence is that of a Treating a vascular disease, particularly atherosclerosis, thrombosis, restenosis, stent restenosis or angioplasty restenosis, by administering a Toll-like receptor-4 (TLR-4) inhibitor to a mammal. This invention relates to a novel method for the treatment of a vascular recognise an epitope containing a part of a domain with not less than 8 amino acids in human CD14 in the region from positions 269-315 in a ful defined sequence of 356 amino acids as given in the specification. The antibody is useful in drug compositions for treating sepsis and for screening remedies for sepsis. This polynucleotide sequence represents anti-CD14 related oligonucleotide of the invention Gaps vascular disease; Toll-like receptor-4 inhibitor; TLR-4 inhibitor; atherosclerosis; restenosis; inflammation; vasotropic; antiarteriosclerotic; thrombolytic; cardiant; antiinflammatory; antisense therapy; gene therapy; transplant atherosclerosis; vein-graft atherosclerosis; thrombosis; stent restenosis; angioplasty restenosis; thrombosis; stent restenosis; reverse transcription polymerase chain reaction; RT-PCR; human; ss; TLR-4; Toll-like receptor 4. ö Match 0.5%; Score 20; DB 1; Length 20; Local Similarity 100.0%; Pred. No. 73; les 20; Conservative 0; Mismatches 0; Indels Human toll-like receptor (TLR) -4 RT-PCR primer Seg ID3. Sequence 20 BP; 4 A; 8 C; 3 G; 5 T; 0 U; 0 Other; Arditi M, Rajavashisth T, Shah PK; (CEDA-) CEDARS SINAI MEDICAL CENT. Example 6; Page 10; 21pp; English. 501 CCCATCCAGAGTTTAGCCCT 520 1 cccarccagagriragcccr 20 BP 23-APR-2002; 2002US-00128166. 24-OCT-2001; 2001US-0335637P. 17-DEC-2001; 2001US-0341359P. ADB39124 standard; DNA; 20 (first entry) WPI; 2003-615988/58. US2003077279-A1. Homo sapiens. 04-DEC-2003 24-APR-2003 ADB39124; Query Match Matches RESULT 65 ADB39124 8888888 ద 

73;

Pred. No.

100.0%;

Best Local Similarity

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      The invention comprises therapeutic agents for preventing or treating diseases associated with endothelial disorders, the agents contain a 3-amino-1,2-benzisothiazole compound. The therapeutic agents of the invention are useful for preventing/treating: diseases associated with endothelial disorders, toxaemia, severe toxaemia, toxic shock haemorrhagic shock, alcohol induced cirrhosis, adult respiratory distress syndrome, chronic rheumatoid arthritis, ulcerative gastritis, Crohn's disease, glomerulonephritis, infectious carditis, systemic lupus erythematosus, seleroderma, Sjoegren's syndrome, multiple organ failure or autoimmune diseases (e.g. multiple sclerosis). The present DNA sequence represents a PCR primer that was used in the exemplification of
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Agents for treating diseases associated with endothelial disorders, toxemia or toll like receptor signaling comprise new or known 3-amino-1,2 -benzisothiazole compounds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           severe toxaemia; toxic shock; haemorrhagic shock; alcohol induced cirrhosis; adult respiratory distress syndrome; chronic rheumatoid arthritis; ulcerative gastritis; Crohn's disease; glomerulonephritis; infectious carditis; systemic lupus erythematosus; scleroderma; Sjoegren's syndrome; multiple organ failure; autoimmune disease; multiple sclerosis; PCR; primer; ss; human;
                                                                                                                                                        Gaps
PCR primer which was used for reverse transcription polymerase chain reaction amplification of human TLR-4 in the exemplification of the
                                                                                                                                                      ..
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         therapeutic agent; endothelial disorder; 3-amino-1; 2-benzisothiazole compound; endothelial disorders; toxaemia;
                                                                                                         0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 73;
tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Mori S;
                                                                                                                                                                                                                                                                                                                                                                                                                                        Human toll-like receptor 4 (TLR4) gene PCR primer #2.
                                                                         Sequence 20 BP; 5 A; 3 C; 4 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 20 BP; 4 A; 8 C; 3 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Mizuno M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Example 4; SEQ ID NO 5; 180pp; Japanese.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Furusako S, Satoh T, Nakamura M,
                                                                                                                                                                                          1768 IGGATACGTTTCCTTATAAG 1787
                                                                                                                                                                                                                             1 regaracerricerrarase 20
                                                                                                                                                                                                                                                                                                                               琚.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     31-MAR-2003; 2003WO-JP004108.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          29-MAR-2002; 2002JP-00132121.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (MOCH ) MOCHIDA PHARM CO LID
                                                                                                                                                                                                                                                                                                                           ADF17208 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                  12-FEB-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  toll-like receptor 4; TLR4
                                                                                                                                Local Similarity 100.
les 20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2003-903121/82
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WO2003087072-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 23-OCT-2003.
                                     invention
                                                                                                                                                                                                                                                                                                                                                                ADF17208;
                                                                                                                Query Match
                                                                                                                                                  Matches
                                                                                                                                                                                                                                                                                     RESULT 66
 888888
                                                                                                                                                                                          ઠ
                                                                                                                                                                                                                          셤
```

DB 1; Length 20;

0.5%; Score 20;

Query Match

```
The invention relates to a system for inhibiting the biological activity of myeloid differentiation factor 88 (MyD88) which comprises an intravascular device and a therapeutic composition coated upon the intravascular device, the composition comprising a MyD88 inhibitor. The system or method is useful for treating vascular diseases including attenced tenosis, transplant atherosclerosis, veln-graft atherosclerosis, thrombosis, restenosis, start restenosis or angioplasty restenosis. It can be used for treating patients suffering from angina pectoris, ischaemias, conditions associated with ischaemias including stroke, transfent schaemic attacks, heart attack, osteonecrosis, colitis, poor kidney function or congestive heart failure, poor blood circulation to the extremities and complications of poor blood circulation including slow wound healing, inflammation, infections and claudication. The present sequence represents a human Toll-like receptor 4 TLR-4 reverse
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               A system for inhibiting the biological activity of myeloid differentiation factor 88 (MyD88), useful for treating vascular diseases (e.g. atherosclerosis), comprises an intravascular device and MyD88
                                                                                                                                                                                                                                                                                                                     myeloid differentiation factor 88; MyD88; atherosclerosis; thrombosis; restenosis; angine pectoris; ischaemia; stroke; heart attack; osteonecrosis; colitis; poor kidney function; congestive heart failure; poor blood circulation; slow wound healing; inflammation; infection; claudication; Toll-like receptor 4; TLR-4; human.
Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match 0.5%; Score 20; DB 1; Length 20; Best Local Similarity 100.0%; Pred. No. 73; Matches 20; Conservative 0; Mismatches 0; Indels
Indels
                                                                                                                                                                                                                                                                                   Human Toll-like receptor 4 TLR-4 RT-PCR primer #1.
.
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 20 BP; 5 A; 3 C; 4 G; 8 T; 0 U; 0 Other;
Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Example 6; SEQ ID NO 3; 18pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Arditi M, Rajavashisth T, Shah PK,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1768 IGGATACGITICCITATAAG 1787
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (CEDA-) CEDARS SINAI MEDICAL CENT.
                                   520
                                                                       20
;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         inhibitor coated on the device.
                                   501 CCCATCCAGAGTTTAGCCCT
                                                           1 CCCATCCAGAGTTTAGCCCT
                                                                                                                                                                    ADI53114 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ranscriptase (RT) - PCR primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         12-DEC-2002; 2002US-00317992.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             17-DEC-2001; 2001US-0341359P.
23-APR-2002; 2002US-00128166.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (first entry)
20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WPI; 2003-897598/82.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              US2003148986-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens.
                                                                                                                                                                                                                                              22-APR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   07-AUG-2003.
Matches
                                                                                                                                RESULT 67
                                                                                                                                                    ADI53114
                                   ð
                                                                     셤
                                                                                                                                                                                      ઠે
```

```
This chimeric phosphorothioate oligonucleotide, having 2'-MOE wings and a
                                                                                                                                                                                                                                                                                                                                                                                                                                      шSc
                                                                                                                                                                                                                                                                                                                                                                                                                                                    /*tag= f
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                'mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                       mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                               mod_base= m5c
                                                                                                                                    20 ccgaaaggrearreres
                                                                                                                                                                            BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            .9-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                                                                                                                                              /*tag= e
/mod_base= 1
                                                                                                                                                                          ACC83594 standard; DNA; 20
                                                                                                                                                                                                                                                                                                  ಹ
                                                                                                                                                                                                           (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                       ъ
                                                                                     Query Match 0.5
Best Local Similarity 100.
Matches 20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                        *tag=
                                                                                                                                                                                                                                                                                         . 20
                                                                                                                                                                                                                                                                                                                                                         *tag=
                                                                                                                                                                                                                                                                                                                                                                                *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WO2003044163-A2
                                                                                                                                                                                                                                                                                  Key
modified_base
                                                                                                                                                                                                                                                                                                                                                modified base
                                                                                                                                                                                                                                                                                                                                                                      modified_base
                                                                                                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                                                       modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                              modified_base
                                                                                                                                                                                                                                                                  Homo sapiens
                                                                                                                                                                                                           08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Karras JG,
                                                                                                                                                                                           ACC83594;
                                                                                                                                                            RESULT 69
                                                                                                                                                                    ACC83594/
8888888888
                                                                                                                                                                                   ò
                                                                                                                                   셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    . .
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The present sequence is that of antisense oligonucleotide ISIS #114646. .
                                                                                                   Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88
                                                                                                                                                                          /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     e.g.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      antisense oligonucleotides for modulating Toll-like receptor 4 gene
                                                                                    Human Toll-like receptor 4 antisense oligonucleotide ISIS #114646
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     expression, particularly useful for preventing, delaying or treat:
inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                            Location/Qualifiers
                                                                                                                                                           /*tag= a
/mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Claim 3; Page 95; 110pp; English.
1 TGGATACGTTTCCTTATAAG 20
                                                                                                                                                                                                                  *tag= b
mod_base= m5c
                                                                                                                                                                                                                                                                                                mod_base= m5c
                                                                                                                                                                                                                                                   mod_base= m5c
                                                                                                                                                                                                                                                                         mod_base= m5c
                                                                                                                                                                                                                                                                                                                         mod_base= m5c
                                                                                                                                                                                                                                                                                                                                        /*tag= g
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                         mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                               /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                              14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                             19-NOV-2001; 2001US-00001863
                               ACC83590/c
ID ACC83590 standard; DNA; 20
                                                                                                                                                                                                                                           υ
                                                                                                                                                                                                                                                                  b
                                                                                                                                                                                                                                                                                                                                                                ч
                                                                                                                                                                                                                                                                                          Φ
                                                                                                                                                                                                                                                                                                                 *tag=
                                                                                                                                                                                                                                                                                                                                                                *tag=
                                                                                                                                                                                                                                                                                                                                                                                       *tag=
                                                                                                                                                                                                                                                                                         *tag=
                                                                                                                                                                                                                                          *tag=
                                                                                                                                                                                                                                                                  *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                              WO2003044163-A2
                                                                                                                                           Key
modified_base
                                                                                                                                                                                                          modified_base
                                                                                                                                                                                                                                                                                                       modified_base
                                                                                                                                                                                                                                                                                                                                                                              modified_base
                                                                                                                                                                                                                                  modified base
                                                                                                                                                                                                                                                         modified_base
                                                                                                                                                                                                                                                                                 modified_base
                                                                                                                                                                                                                                                                                                                                modified_base
                                                                                                                                                                                                                                                                                                                                                       modified base
                                                                                                                            Homo sapiens
                                                                     08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     responses
                                                      ACC83590;
                       RESULT
셤
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
deoxy gap, is targeted to the coding region of human Toll-like receptor 4 man. The exhibite 85% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /mcd base= OTHER /mcdbase= OTHER = phosphorothioate nucleotides, the /note= "OTHER = phosphorothioate nucleotides, the oligomucieotide comprises a central gap region of 10 2'-deoxymucieotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Human Toll-like receptor 4 antisense oligonucleotide ISIS #114650
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                   0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 73;
ive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 6 A; 8 C; 2 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2354 CCGAAAGGTGATTGTTGTGG 2373
```

```
The present sequence is that of antisense oligonucleotide ISIS #114650. This chimeric phosphorothicate oligonucleotide, having 2'-MOE wings and a decxy ago, is targeted to the 3' untranslated region of human Toll-like receptor 4 mRNA. It exhibits 79% inhibition of human Toll-like receptor expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Th1 or Th2 responses
New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                receptor 4; receptor; antiinflammatory; immunomodulator;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          /mod_base= OTHER
/note= "OTHER = phosphorothicate nuclectides, the
oligonuclectide comprises a central gap region of 10 2'-
deoxynuclectides, flanked on both sites by 5-nuclectides
wings composed of 2'-methoxyethyl nuclectides"
                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Human Toll-like receptor 4 antisense oligonucleotide ISIS #114629
                                                                                                                                                                                                                                                                                             ;
                                                                                                                                                                                                                                                           DB 1; Length 20; 73;
                                                                                                                                                                                                                                                                                             0; Indels
                                                                                                                                                                                                                                       Sequence 20 BP; 7 A; 5 C; 5 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                         0.5%; bccc.
100.0%; Pred. No. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Location/Qualifiers
                                                                                                                                                                                                                                                                                                                         2884 TCTTCCAGGTGGGCATTTCA 2903
                                                                      Claim 3; Page 95; 110pp; English
                                                                                                                                                                                                                                                                                                                                         20 TCTTCCAGGTGGGCATTTCA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  phosphorothioate; antisense; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      m50
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              mSG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          /*tag= e
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                        BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      14-NOV-2002; 2002WO-US036390
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /*tag= d
/mod_base= 1
                                                                                                                                                                                                                                                                                                                                                                                                             .606/c
ACC83606 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      mod_base=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ಗ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Ω
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          υ
                                                                                                                                                                                                                                                                                             20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       *tag=
                                                                                                                                                                                                                                                                               Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human; Toll-like
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WO2003044163-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            30-MAY-2003
                                            responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACC83606;
                                                                                                                                                                                                                                                                   Query Match
                                                                                                                                                                                                                                                                                            Matches
                                                                                                                                                                                                                                                                                                                                                                                           RESULT 70
ACC83606/
                                                                                                                                                                                                                                                                                                                                                                                                                        à
                                                                                                                                                                                                                                                                                                                                               셤
```

(ISIS-) ISIS PHARM INC

```
ö
                                                                                                                                                                                                                                                          associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Th1 or Th2 responses
                                                           New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; ss.
                                                                                                                                                        The present sequence is that of antisense oligonucleotide ISIS #114629. This chimeric phosphorothioate oligonucleotide, having 2'-WOB wings and selecty gap, is trageted to the S' untranslated region of human Toll-like receptor 4 mRNA. It exhibits 48% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human Toll-like receptor 4 antisense oligonucleotide ISIS #114638
                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                              0.5%; Score 20; DB 1; Length 20,
                                                                                                                                                                                                                                                                                                                                                         Indels
                                                                                                                                                                                                                                                                                                      Seguence 20 BP; 7 A; 4 C; 6 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                           .
0
                                                                                                                                                                                                                                                                                                                                        100.0%; Pred. ...
                                                                                                                                   Example 14; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1. .20
/*tag= a
/mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                     225 CTTCTTGCTAAATGCTGCCG 244
                                                                                                                                                                                                                                                                                                                                                                                                               Н
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /*tag= e
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                      20 CTTCTTGCTAAATGCTGCG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ACC83611 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     υ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ø
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           *tag= f
                                                                                                                                                                                                                                                                                                                                                           20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            *tag=
           Karras JG, Koller E;
                                      WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                            Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO2003044163-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           08-SEP-2003
                                                                                                            responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ACC83611;
                                                                                                                                                                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                           Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 71
ACC83611/
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 à
                                                                                                                                                                                                                                                                                                                                                                                                               g
                                                                                                                                                                                                                                                                                         ö
```

```
ACC83583;
                                                                                                                                                                                                                                                                                                                                                                                                                                        Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ACC83583/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 73
  셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                              The present sequence is that of antisense oligonuclectide ISIS #114638. This chimeric phosphorothioate oligonucleotide, having 2'-MOB wings and a deoxy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 11% inhibition of human Toll-like receptor 4 expression in THP-1 cells. More active oligonucleotides are useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                          New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /note= "OTHER = phosphorothioate nucleotides, the oligomucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Human Toll-like receptor 4 antisense oligonucleotide ISIS #114637
                                                                                                                                                                                                                                                                                                                                                 ;
0
                                                                                                                                                                                                                                                                                                                             DB 1; Length 20; 73;
                                                                                                                                                                                                                                                                                                                                                 0; Indels
                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 5 A; 6 C; 3 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                            0.5%; Score 20; DB 100.0%; Pred. No. 73; ative 0; Mismatches
                                                                                                                                                                              Example 14; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                    1586 AGTCTTGAAATGGCTGGCA 1605
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                        20 AGTCTTGAAATGGCTGGCA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            *tag= c
mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                        ACC83584 standard; DNA; 20 BP
                            14-NOV-2002; 2002WO-US036390.
                                               19-NOV-2001; 2001US-00001863
                                                                                                                                                                                                                                                                                                                 ъ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              08-SEP-2003 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        *tag= b
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     /*tag=
                                                                  (ISIS-) ISIS PHARM INC
                                                                                                        WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
         30-MAY-2003.
                                                                                                                                                          responses.
                                                                                                                                                                                                                                                                                                                                                                                                                                                            ACC83584';
                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 72
ò
                                                                                                                                                                                                                                                                                                                                                                                     셤
```

```
The present sequence is that of antisense oligonucleotide ISIS #114637. This chimeric phosphorothioate oligonucleotide, having 2'-MOE wings and a deoxy gap, is targeted to the coding region of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibition the expression of Toll-like receptor 4 in cells or itssues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Human, Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Human Toll-like receptor 4 antisense oligonucleotide ISIS #114633
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /note= "OTHER = phosphorothioate nucleotides, the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.5%; Score 20; DB 1; Length 20; ilarity 100.0%; Pred. No. 73; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Sequence 20 BP; 5 A; 6 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       998 GGCCTGTGCAATTTGACCA 1017
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1. .20
/*tag= a
/mod_base= OTHER
mod_base= m5c
                                                                                      /mod_base= m5c
                                                                                                                                                                                                                                       /*tag= g
/mod_base= m5c
                                                                                                                                                                            'mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20 ééécérérécsarrreacea
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                                                                                                                                                               14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACC83583 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (first entry)
                                                                                                                                                *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (ISIS-) ISIS PHARM INC.
                                                             *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Karras JG, Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Best Local Similarity
Matches 20; Conserva
                                                                                                                                                                                                                                                                                                                             WO2003044163-A2
                              modified_base
                                                                                                                 modified base
                                                                                                                                                                                                        modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                       30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          responses
```

Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothicate; antisense; 88.

Location/Qualifiers

ಡ

. .20 \*tag=

Key modified\_base

Homo sapiens

Human Toll-like receptor 4 antisense oligonucleotide ISIS #114648.

08-SEP-2003 (first entry)

```
The present sequence is that of antisense oligonucleotide ISIS #114633. This chimeric phosphorothicate oligonucleotide, having 2'-MOB wings and a decoxy gap, is trargeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 69% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               New antisense oligonuclectides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; ive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 6 A; 7 C; 6 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                          /*tag= e
mod_base= m5c
                                                                                                                                                                                                                                                                                                                                     /mod_base= m5c
                                                                                                                  mod_base= m5c
                                                                                                                                                *tag= d
mod_base= m5c
                                                                                                                                                                                                                                  '*tag= f
'mod_base= m5c
                                                                                                                                                                                                                                                                                           'mod_base= m5c
                                                            *tag= b
mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                        14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                    19-NOV-2001; 2001US-00001863
                                                                                                                                                                                                                                                                               þ
                                                                                                                                                                                                                                                                                                                        Ø
                                                                                                                                              ø
                                                                                                      *tag=
                                                                                                                                                                                                                                                                                                                        *tag=
                                                                                                                                                                                                                                                                              *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
Best Local Similarity
Matches 20; Conserv
                                                                                                                                                                                                                                                                                                                                                                  WO2003044163-A2
                                                                                                                                                                           modified base
                                                                                                                                                                                                                     modified_base
                                                                                                                                                                                                                                                                                                       modified base
                                             modified base
                                                                                       modified_base
                                                                                                                                  modified_base
                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                          30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                responses.
```

```
The present sequence is that of antisense oligonucleotide ISIS #114648. This chimeric phosphorothioate oligonucleotide, having 2'-MOB wings and a decoyy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 73% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
                                                                                                                                        /mod=base= OTHER
/note= "OTHER = phosphorothioate nucleotides, the
/note= "OTHER = phosphorothioate nucleotides of 10 2'-
oligonucleotide comprises a central gap region of 10 2'-
deoxynucleotides, flanked on both sites by 5-nucleotides
wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                                                        /*tag= b
/mod_base= m5c
                                                                                                                                                                                                                                                                           mSc
                                                                                                                                                                                                                                                                                                                mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                      mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                /*tag= g
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /*tag= i
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                  mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                              /*tag= c
/mod_base= 1
                                                                                                                                                                                                                                                                                                                                         Ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                     *tag= h
                                                                                                                                                                                                                                                                                                   D
                                                                                                                                                                                                                                                                                                                                                                              *tag=
                                                                                                                                                                                                                                                                                                   *tag=
                                                                                                                                                                                                                                                                                                                                         *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WO2003044163-A2
                                                                                                                                                                                                             modified base
                                                                                                                                                                                                                                                   modified base
                                                                                                                                                                                                                                                                                       modified base
                                                                                                                                                                                                                                                                                                                            modified_base
                                                                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                                    modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                          modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   responses.
```

ö

Gaps

ö

516 GCCCTGGGAGCCTTTTCTGG 535

20; Conservative

GCCTGGGAGCCTTTTCTGG

20

a ò

RESULT 74
ACC83592/c
ID ACC83592 standard; DNA; 20 BP.
XX
XA ACC83592;
XX

Wed Feb 16 09:29:34 2005

10001863-3.81.rng

```
WO2003044163-A2
                                                                                                                                                                                                                                                                                                                                                                      Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            modified base
                                                                                                                                                                                                                                                              08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                Ното варієпв
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 30-MAY-2003
                                                                                                                                                                                                                                       ACC83608;
                                                                                                                                                                                                    ACC83608/
                                                                                                                                                                                        RESULT
  ន្តដ្ឋប្បន្នប្ល
                                                                                                                               ð
                                                                                                                                                    d
                                              ö
                                                                                                                                                                                                                                                Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                      /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             e.9
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             The present sequence is that of antisense oligonucleotide ISIS #114628. This chimeric phosphorothicate oligonucleotide, having 2'-MOE wings and adeoxy gap, is targeted to the 5' untranslated region of human 7011-like receptor 4 mRNA. It exhibits 33 inhibition of human 7011-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of
                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           expression, particularly useful for preventing, delaying or treating inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              New antisense oligonucleotides for modulating Toll-like receptor 4
                                                                                                                                                                                                                         Human Toll-like receptor 4 antisense oligonucleotide ISIS #114628
                                             ;
                     DB 1; Length 20; 73;
                                             Indels
Sequence 20 BP; 4 A; 8 C; 5 G; 3 T; 0 U; 0 Other;
                                             ö
             0.5%; Scor.
100.0%; Pred. No. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Example 14; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                            Location/Qualifiers
                                                                   GGGCGCCACATCTTCTGGAG 2590
                                                                                                                                                                                                                                                                                                                                             mod_base= OTHER
                                                                                 20 GGGCGCACATCTTCTGGAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                   /*tag= b
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                      *tag= c
mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         /*tag= d
'mod_base= m5c
                                                                                                                                                     ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  19-NOV-2001; 2001US-00001863.
                                                                                                                                         ACC83605/c
ID ACC83605 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ъ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          *tag= e
                                             Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2003-468766/44.
                                 Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WO2003044163-A2
                                                                                                                                                                                                                                                                                                            Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                       modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                        modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                   Ното варіепв
                                                                                                                                                                                                 08-SEP-2003
                                            20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Karras JG,
                                                                   2571
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   responses
                                                                                                                                                                          ACC83605;
                     Query Match
                                            Matches
                                                                                                                              RESULT
                                                                                        ద
                                                                                                                                                     ઠે
```

```
ö
Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition a associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Th1 or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                   Human, Toll-like receptor 4; receptor; antiinflammatory; immunomodulator;
phosphorothioate; antisense; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                             Gape
                                                                                                                                                                                                                                                                                                                                         Human Toll-like receptor 4 antisense oligonucleotide ISIS #114634.
                                                                                                                            ö
                                                                                            ch 0.5%; Score 20; DB 1; Length 20; 1 Similarity 100.0%; Pred. No. 73; 20; Conservative 0; Mismatches 0; Indels
                                                                    Sequence 20 BP; 3 A; 4 C; 10 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
                                                                                                                                                      193 CGGAGCCTCAGCCTTCACC 212
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 mod_base= OTHER
                                                                                                                                                                      20 CGGAGCCTCAGCCCTTCACC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /*tag= c
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /*tag= d
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              /*tag= e
'mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     /*tag= f
/mod_base= m5c
                                                                                                                                                                                                                                                       BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                     ACC83608 standard, DNA; 20
                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ಹ
                                                                                                                                                                                                                                                                                                                                                                                                                                                       . .20
*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Karras JG, Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2003-468766/44.
                                                                                             Query Match
Best Local Similarity
Matches 20; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          responses.
```

```
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          g
                                                                                                                                                                                                                                                                                                                                                                       ö
                                                   The present sequence is that of antisense oligonucleotide ISIS #114634. This chimeric phosphorothicate oligonucleotide, having 2'-MOE wings and a decoxy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibites 55% inhibition of human Toll-like receptor 4 expression in THP-1 cells. Such oligonucleotides are useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /notes "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               New antisense oligonucleotides for modulating Toll-like receptor 4 gene
                                                                                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human Toll-like receptor 4 antisense oligonucleotide ISIS #114635
                                                                                                                                                                                                                                                                                                                                                                       ..
                                                                                                                                                                                                                                                                                                                        Query Match 0.5%; Score 20; DB 1; Length 20; Best Local Similarity 100.0%; Pred. No. 73; Matches 20; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                      Sequence 20 BP; 6 A; 6 C; 6 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Location/Qualifiers
                       Example 14; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                518 CCTGGGAGCCTTTCTGGAC 537
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                      20 ccreedaaccrirircreaac 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /mod base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             踣.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                14-NOV-2002; 2002WO-US036390
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       19-NOV-2001; 2001US-00001863
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ACC83609 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ಹ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Д
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WO2003044163-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ACC83609
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 77
ACC83609/c
LD ACC83609/c
SXX
XX
XX
XX
DB Humar
XXX
CS Homo
XX
CS Ho
      ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                      g
```

```
·,
                                                                                                                     The present sequence is that of antisense oligonucleotide ISIS #114635. This chimeric phosphorothioate oligonucleotide, having 2'-MOE wings and a deoxy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 51% inhibition of human Toll-like receptor 4 expression in THP-1 cells. Such oligonucleotides are useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /note= "OTHER = phosphorothioate nucleotides, the oligomucleotide comprises a central gap region of 10 2'-deoxymucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
      e.g.
delaying or treating
Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human Toll-like receptor 4 antisense oligonucleotide ISIS #114652
                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                          Score 20; DB 1; Length 20;
Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                  0; Indels
                                                                                                                                                                                                                                                                                                                                                              Sequence 20 BP; 5 A; 3 C; 8 G; 4 T; 0 U; 0 Other;
    expression, particularly useful for preventing, inflammatory disorders, or conditions involving
                                                                                                                                                                                                                                                                                                                                                                                               0.5%; Scor.
100.0%; Pred. No. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
                                                                                        Example 14; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              791 CCTGTCCCTGAACCCTATGA 810
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      'mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /*tag= h
/mod_base= m5c
20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            щSС
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    шSc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  mSc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 mSc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20 CCTGTCCCTGAACCCTATGA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /*tag= c
/mod_base= n
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /*tag= d
/mod_base= 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /*tag= e
/mod_base= 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    /*tag= g
/mod_base= 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            mod_base=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ACC83596 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                         20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ...20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  08-SEP-2003
                                                       responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACC83596;
                                                                                                                                                                                                                                                                                                                                                                                                                                                           Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ACC83596/
```

```
Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                         30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ACC83588;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
Best Local S
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 80
      g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                        New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Th1 or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         The present sequence is that of antisense oligonucleotide ISIS #114652. This chimeric phosphorothioate oligonucleotide, having 2'-MOB wings and edeoxy gap, is targeted to the 3' untranslated region of human Toll-like receptor 4 mRNA. It exhibits 85% inhibition of human Toll-like receptor expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human Toll-like receptor 4 antisense oligonucleotide ISIS #114626
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 20 BP; 4 A; 8 C; 4 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       100.0%; Pred. No. 73; tive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       3144 GGGCTCCTGATGCAAGATG 3163
                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Claim 3; Page 95; 110pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /mod base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 GGGGCTCCTGATGCAAGATG 1
                   /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ACC83603 standard; DNA; 20 BP
                                                                                                                                              14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                        19-NOV-2001; 2001US-00001863
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         08-SEP-2003 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     /*tag= ,c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          *tag= b
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ...20
/*tag=
                                                                                                                                                                                                                                  (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                           Koller E;
                                                                                                                                                                                                                                                                                                                 WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Best Local Similarity
                                                             WO2003044163-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens
                                                                                                       30-MAY-2003
                                                                                                                                                                                                                                                                       Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                               responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ACC83603;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RESULT 79
ACC83603/
THE LITER FOR SON THE STATE OF 
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ઠે
```

```
The present sequence is that of antisense oligonucleotide ISIS #114626. This chimeric phosphorothioate oligonucleotide, having 2'-MOB wings and a decoxy gap, is targeted to the 5' untranslated region of human Toll-like receptor 4 mRNA. It exhibits 15% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 New antisense oligonuclectides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human Toll-like receptor 4 antisense oligonucleotide ISIS #114643.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Score 20; DB 1; Length 20;
Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Sequence 20 BP; 5 A; 7 C; 6 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Example 14; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 79
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 phosphorothioate; antisense; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20 CGCCTGGCTGGGACTCTGAT 1
                                                                                                                                                                                                                                                                                       /reg= g
/mod_base= m5c
                                                                        mod_base= m5c
                                                                                                                             /*tag= e
/mod_base= m5c
                                                                                                                                                                                                                               mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                     /*tag= h
/mod_base= m5c
mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.5%; S
100.0%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              60 CGCCTGGCTGGGACTCTGAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        19-NOV-2001; 2001US-00001863
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ACC83588/c
ID ACC83588 standard; DNA; 20
                                                  T
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20; Conservative
                                                                                                                                                                                                                 *tag=
                                                  *tag≂
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Karras JG, Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                         WO2003044163-A2
                      modified_base
                                                                                                    modified_base
                                                                                                                                                                               modified_base
                                                                                                                                                                                                                                                                modified_base
                                                                                                                                                                                                                                                                                                                                              modified base
```

Human, Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; ss.

Homo sapiens

Human Toll-like receptor 4 antisense oligonucleotide ISIS #114653

(first entry)

08-SEP-2003

ACC83597;

```
The present sequence is that of antisense oligonucleotide ISIS #114643. This chimeric phosphorothicate oligonucleotide, having 2'-MOE wings and a decoyy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 86% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                  /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 73;
tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 20 BP; 8 A; 6 C; 5 G; 1 T; 0 U; 0 Other;
           .ocation/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Claim 3; Page 95; 110pp; English.
                                                      mod base= OTHER
                                                                                                                                                    'mod_base= m5c
                                                                                                                                                                                                                                                                                  mod_base= m5c
                                                                                                                                                                                                                                                                                                             /*tag= f
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                      /mod_base= m5c
                                                                                                                                                                                                mod_base≈ m5c
                                                                                                                                                                                                                                        mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                        14-NOV-2002; 2002WO-US036390
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   19-NOV-2001; 2001US-00001863
                                                                                                                                        Д
                                                                                                                                                                                                                                                                                                                                                         מ
                                                                                                                                                                                                                          ש
                                                                                                                                                                                *tag=
                                                                                                                                                                                                                          *tag=
                                                                                                                                                                                                                                                                                                                                                        *tag=
                                                                                                                                         *tag=
                                                                                                                                                                                                                                                                  *tag=
                                      *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                 WO2003044163-A2
              Key
modified base
                                                                                                                                                                   modified_base
                                                                                                                                                                                                           modified_base
                                                                                                                                                                                                                                                      modified_base
                                                                                                                           modified base
                                                                                                                                                                                                                                                                                              modified_base
                                                                                                                                                                                                                                                                                                                                       modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                            30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             responses.
```

```
condition involving an immune response, particularly Th1 or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   The present sequence is that of antisense oligonucleotide ISIS #114653. This chimeric phosphorothicate oligonucleotide, having 2. *MDE wings and a decay gap, is trageted to the 3' untranslated region of human Toll-like receptor 4 mRNA. It exhibits 72% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a
                                                                                                                                                                                       /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              BP; 7 A; 6 C; 3 G; 4 T; 0 U; 0 Other;
                                                                                                                                            Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                OTHER
                                                                                                                                                                                                                                                   /*tag= b
/mod_base= m5c
                                                                                                                                                                                                                                                                                                        mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                               mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                   mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                          /*tag= g
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                             mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     19-NOV-2001; 2001US-00001863.
                                                                                                                                                         1. .20
/*tag= a
/mod_base= (
                                                                                                                                                                                                                                                                                            υ
                                                                                                                                                                                                                                                                                                                                                                    Φ
                                                                                                                                                                                                                                                                                                                                                                                                        *tag=
                                                                                                                                                                                                                                                                                          *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (ISIS-) ISIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                *tag=
                                                                                                                                                                                                                                                                                                                                                                    *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Karras JG, Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WO2003044163-A2
                                                                                                                                                         modified base
                                                                                                                                                                                                                                            modified base
                                                                                                                                                                                                                                                                                modified base
                                                                                                                                                                                                                                                                                                                   modified_base
                                                                                                                                                                                                                                                                                                                                                       modified base
                                                                                                                                                                                                                                                                                                                                                                                           modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                               modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          responses
```

DB 1; Length 20;

0.5%; Score 20;

Query Match

ô

Gaps

ö

2137 TGATGCTTCTTGCTGGCTGC 2156

20; Conservative

Matches

ò

Query Match Best Local Similarity

rgargcrrcrrgcrggcrgc

20

ACC83597 standard; DNA; 20 BP.

RESULT 81 ACC83597/c ID ACC83

셤 Š

```
ö
                                         This chimeric phosphorothioate oligonucleotide, having 2'-MOE wings and a decoxy gap, is targeted to the 5' untranslated region of human Toll-like receptor 4 mRNA. It exhibits 47% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is a particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human, Toll-like receptor 4; receptor; antiinflammatory; immunomodulator;
phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                            present sequence is that of antisense oligonucleotide ISIS #114625
                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Human Toll-like receptor 4 antisense oligonucleotide ISIS #114627,
                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                            DB 1; Length 20; 73;
                                                                                                                                                                                                                                                                                  0; Indels
                                                                                                                                                                                                          Sequence 20 BP; 5 A; 7 C; 7 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                            Score 20; DB 1; Pred. No. 73; 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
                                                                                                                                                                                                                                 0.5%; Scc.
100.0%; Prev
0; V
                                                                                                                                                                                                                                                                                                                      26
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           'mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                      resescrescresearer 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /*tag= c
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /*tag= g
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                mod_base= m5c
                                                                                                                                                                                                                                                                                                                    57 TCGCGCCTGGCTGGGACTCT
                                                                                                                                                                                                                                                                                                                                                                                                                                                   ВP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                    604/c
ACC83604 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Ø
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ರ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             44
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       *tag= h
                                                                                                                                                                                                                                                                                20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ...20
*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               *tag=
                                                                                                                                                                                                                                              Query Match
Best Local Similarity
Matches 20; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WO2003044163-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        domo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ACC83604;
                                                                                                                                                                                                                                                                                                                                                        20
                                                                                                                                                                                                                                                                                                                                                                                                                                  ACC83604/
                                                                                                                                                                                                                                                                                                                                                                                                               RESULT
           ઠે
                                                                                                                                                                                                                                                                                                                                                      셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                     Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator;
phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    /notē= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxymucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                            Gaps
                                                                                                                                                                                                                                                                                                 Human Toll-like receptor 4 antisense oligonucleotide ISIS #114625.
                            ö
                            Indels
                            ö
100.0%; Prec. ...
         Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Example 14; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                              location/Qualifiers
                                                            3521 TIGITTAAAGGGGGCACTCT 3540
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     base= OTHER
                                                                                              20 rrgrrraaaccccccrcr 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                '*tag= e
mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             mod_base= m5c
                                                                                                                                                                                         ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                         ACC83602 standard; DNA; 20
                                                                                                                                                                                                                                                               (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      đ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        O
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             þ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         b
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               *tag= h
                          20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            'tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2003-468766/44.
   Best Local Similarity
Matches 20; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO2003044163-A2
                                                                                                                                                                                                                                                                                                                                                                                                                              Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            modified base
                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens
                                                                                                                                                                                                                                                             08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      responses
                                                                                                                                                                                                                           ACC83602;
                                                                                                                                                   RESULT 82
ACC83602(XX
XX
ACC83602(XX
XX
ACC83602(XX
ACC830-AC
ACC83602(XX
ACC8
```

```
Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         modified base
                                                                                               19-NOV-2001;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             08-SEP-2003
                30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20;
                                                                                                                                                                                                                                                                                                                         responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ACC83589;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Best Loc
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 85
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ACC83589/
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                              The present sequence is that of antisense oligonucleotide ISIS #114627. This chimeric phosphorothicate oligonucleotide, having 2'-MOE wings and a deoxy gap, is trageted to the S' untranslated region of human Toll-like receptor 4 mRNA. It exhibits 30% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Poll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /mod_base= OTHER /mosphorothioate nucleotides, the /note= "OTHER = phosphorothioate nucleotides, the oligomucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                       New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human Toll-like receptor 4 antisense oligonucleotide ISIS #114631
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Score 20; DB 1; Length 20;
Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 5 A; 7 C; 6 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             100.0%; Pred. No. 73; ative 0; Mismatches
                                                                                                                                                                                                                                                             Example 14; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         128 CTGCGTGGAGACTTGGCCCT 147
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20 CTGCGTGGAGACTTGGCCCT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /*tag= e
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ВР
19-NOV-2001; 2001US-00001863
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.5%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ACC83581 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ಹ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               08-SEP-2003 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        υ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              V
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Ω
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      *tag=
                                       (ISIS-) ISIS PHARM INC
                                                                                                                  WPI; 2003-468766/44.
                                                                              Koller
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WO2003044163-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Homo sapiens
                                                                              Karras JG,
                                                                                                                                                                                                                         responses.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ACC83581;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 84
ACC83581/
ACC835
ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               셤
```

```
ó
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        The present sequence is that of antisense oligonucleotide ISIS #114631. This chimeric phosphorothioate oligonucleotide, having 2'-MOB wings and a decoyy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 73% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human, Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; ss.
                                                                                                                                                                                                                                                                                                  New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    /mod_base= OTHER
/note= "OTHER = phosphorothicate nuclectides, the
oligonuclectide comprises a central gap region of 10 2'-
deoxynuclectides, flanked on both sites by 5-nuclectides
wings composed of 2'-methoxyethyl nuclectides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Human Toll-like receptor 4 antisense oligonucleotide ISIS #114644.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               .;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Sequence 20 BP; 3 A; 4 C; 6 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        453 GCATATCAGAGCCTAAGCCA 472
                                                                                                                                                                                                                                                                                                                                                                                                                                                      Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Н
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /mod base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    20 gcararcagagccraagcca
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        BP
                                                        2001US-00001863.
14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         /mod_base=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ACC83589 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ಥ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               שי
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          *tag=
                                                                                                                    (ISIS-) ISIS PHARM INC
                                                                                                                                                                                Karras JG, Koller E;
                                                                                                                                                                                                                                         WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Local Similarity
```

```
The present sequence is that of antisense oligonucleotide ISIS #114632. This chimeric phosphorothioate oligonucleotide, having 2'-MOE wings and a decay gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 72% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human Toll-like receptor 4 antisense oligonucleotide ISIS #114649.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Score 20; DB 1; Length 20;
Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 20 BP; 5 A; 3 C; 8 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  501 CCCATCCAGAGTTTAGCCCT 520
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /mod base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Н
                                                                             mod_base= m5c
                                                                                                                   /*tag= c
/mod_base= m5c
                                                                                                                                                                                            /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.5%; S
100.0%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    20 cccarccadacriradcccr
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        BP
                                                                                                                                                                                                                                                                                                                                                19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                            14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ACC83593/c
ID ACC83593 Btandard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (first entry)
                                                                                                                                                                           b
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match 0.5
Best Local Similarity 100.
Matches 20; Conservative
                                                                                                                                                                           tag=
                                                           *tag=
                                                                                                                                                                                                                                                                                                                                                                                     (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                             Karras JG, Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WPI; 2003-468766/44.
                                                                                                                                                                                                                                WO2003044163-A2
                                         modified base
                                                                                             modified_base
                                                                                                                                                       modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          nodified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             08-SEP-2003
                                                                                                                                                                                                                                                                      30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                responses.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACC83593;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 87
     g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      THERES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          The present sequence is that of antisense oligonucleotide ISIS #114644. This chimeric phosphorothioate oligonucleotide, having 2'-MOE wings and a decoyy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 72% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                             New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /noce= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Human Toll-like receptor 4 antisense oligonucleotide ISIS #114632
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 73;
iive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 6 A; 6 C; 6 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           2259 GGGGTGCCTCCATTTCAGCT 2278
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              20 GGGGTGCCTCCATTTCAGCT 1
                     /*tag= e
/mod_base= m5c
                                                                                                                                   /*tag= g
/mod_base= m5c
                                                                                             /mod_base= m5c
                                                                                                                                                                                                                                                                  14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                        19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              582/c
ACC83582 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1. .20
/*tag= a
/mod_base=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (first entry)
                                                                         /*tag= f
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20; Conservative
                                                                                                                                                                                                                                                                                                                                              (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                   Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Local Similarity
                                                                                                                                                                                          WO2003044163-A2
modified base
                                                     modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Key
modified_base
                                                                                                               modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       08-SEP-2003
                                                                                                                                                                                                                            30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                   Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACC83582
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   datches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 86
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ACC83582/
```

ö

ሯ g Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.

Homo sapiens

Human Toll-like receptor 4 antisense oligonucleotide ISIS #114630

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        The present sequence is that of antisense oligonucleotide ISIS #114649. This chimeric phosphorothioate oligonucleotide, having 2'-MOE wings and a decoxy gap, is targeted to the 3' untranslated region of human Toll-like receptor 4 mRNA. It exhibits 91% inhibition of human Toll-like receptor 7 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
/note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Score 20; DB 1; Length 20;
Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Seguence 20 BP; 4 A; 6 C; 4 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       100.0%; Prea. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                            /*tag= e
/mod_base= m5c
                                                                                                                                                                                                                                    /*tag= f
/mod_base= m5c
                                                                                                                                                                                                                                                                                              m5c
                                                                       /*tag= b
/mod_base= m5c
                                                                                                                           /mod_base= m5c
                                                                                                                                                                     mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                             14-NOV-2002; 2002WO-US036390
                                                                                                                                                                                                                                                                                                                                                                                                        19-NOV-2001; 2001US-00001863
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.5%;
                                                                                                                                                                                                                                                                               /*tag= g
/mod_base=
                                                                                                                                                       ש
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20; Conservative
                                                                                                              *tag=
                                                                                                                                                         *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                    (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                             Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Local Similarity
                                                                                                                                                                                                                                                                                                                         WO2003044163-A2
                                                                                                   modified base
                                                                                                                                           modified_base
                                                                                                                                                                                  modified base
                                                                                                                                                                                                                          modified_base
                                                                                                                                                                                                                                                                  modified base
                                                          modified_base
                                                                                                                                                                                                                                                                                                                                                   30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                             Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            responses.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Matches
```

2821 AGGGCTGCTAATCTCAAGGA 2840

AGGGCTGCTAATCTCAAGGA

20

셤 8

BP

(first entry)

08-SEP-2003

ACC83607

ACC83607/c ID ACC83607 standard; DNA; 20

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           The present sequence is that of antisense oligonucleotide ISIS #114630. This chimeric phosphorothioate oligonucleotide, having 2'-MOE wings and a deoxy gap, is targeted to the start codon region of human Toll-like receptor 4 mRNA. It exhibits 48% inhibition of human Toll-like receptor 4 profile in the start of the start of the expression of Toll-like receptor 4. In cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
                                                                                                                                   /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 .
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         DB 1; Length 20; 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 7 A; 5 C; 3 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.5%; Score 20; DB
ilarity 100.0%; Pred. No. 73;
Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Example 14; Page 95; 110pp; English.
                                                                                       Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             282 TGCATGGAGCTGAATTTCTA 301
                                                                                                                            OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      20 recardeadcreaarrrcra 1
                                                                                                                                                                                                        /*tag= b
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                         mSc
                                                                                                                                                                                                                                                                                                                                                                              mSc
                                                                                                                                                                                                                                                                                                  mod_base= m5c
                                                                                                                                                                                                                                              '*tag= c
'mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                                                                                                                                                                            14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                           /*tag= e
/mod_base= 1
                                                                                                                                                                                                                                                                                                                                                                /*tag= f
/mod_base= n
                                                                                                                              mod base=
                                                                                                                 ď
                                                                                                                                                                                                                                                                                     Ъ
                                                                                                                                                                                                                                                                                       *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Karras JG, Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Local Similarity
es 20; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                        WO2003044163-A2
                                                                                         Key
modified_base
                                                                                                                                                                                             modified base
                                                                                                                                                                                                                                   modified_base
                                                                                                                                                                                                                                                                                                                                                      modified base
                                                                                                                                                                                                                                                                          modified_base
                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             SISI (-SISI)
                                                                                                                                                                                                                                                                                                                                                                                                                                   30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Best Loca
Matches
Ś
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      셤
```

10001863-3.81.rng

```
The present sequence is that of antisense oligonucleotide ISIS #114636. This chimeric phosphorothioate oligonucleotide, having 2'-MOE wings and a deoxy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 59% inhibition of human Toll-like receptor 4 expression in THP-1 cells. Such oligonucleotides are useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                     Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; ss.
                                                                                                                                                                                                                                                             /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           New antisense oligonucleotides for modulating Toll-like receptor 4 ge expression, particularly useful for preventing, delaying or treating inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                         Human Toll-like receptor 4 antisense oligonucleotide ISIS #114636.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 20 BP; 8 A; 6 C; 3 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Example 14; Page 95; 110pp; English.
                                                                                                                                                                                                          Location/Qualifiers
                                                                                                                                                                                                                                   /*tag= a
/mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                           '*tag= c
'mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                   mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                     mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     /mod_base= m5c
                            BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               19-NOV-2001; 2001US-00001863.
                           ACC83610 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                     ъ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Φ
                                                                                                                                                                                                                                                                                                                                    *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                     *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WO2003044163-A2
                                                                                                                                                                                                         Key
modified_base
                                                                                                                                                                                                                                                                                                                      modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                 modified_base
                                                                                                                                                                                                                                                                                                                                                                                                       modified base
                                                                               08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                              modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         modified base
                                                                                                                                                                                sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Karras JG,
                                                     ACC83610;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        responses
                                                                                                                                                                                Homo
RESULT 89
```

```
ö
                                                                                                                                                                                         Human, Toll-like receptor 4; receptor; antiinflammatory; immunomodulator;
phosphorothioate; antisense; ss.
                                                                                                                                                                                                                                                                                /note= "OTHER = phosphorothioate mucleotides, the oligomucleotide comprises a central gap region of 10 2'-deoxymucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                     Gaps
                                                                                                                                                                    Human Toll-like receptor 4 antisense oligonucleotide ISIS #114645
                     ö
  Length 20
                    0; Indels
0.5%; Score 20; DB 1;
100.0%; Pred. No. 73;
ive 0; Mismatches
                                                                                                                                                                                                                                          Location/Qualifiers
                                        903 ATTCAAGGICIGGCTGGTTT 922
                                                                                                                                                                                                                                                                        OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                          /*tag= e
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     /*tag= e
mod_base= m5c
                                                                                                                                                                                                                                                                                                                                            mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                          mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                        mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /mod base= m5c
                                                    20 ATTCAAGGTCTGGCTGGTTT
                                                                                                            BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     14-NOV-2002; 2002WO-US036390.
                                                                                                           ACC83613 standard; DNA; 20
                                                                                                                                                                                                                                                                      /mod_base=
                                                                                                                                                  (first entry)
                                                                                                                                                                                                                                                               ซ
                                                                                                                                                                                                                                                                                                                                                                 U
                                                                                                                                                                                                                                                                                                                                                                                             В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /*tag= f
                    Conservative
                                                                                                                                                                                                                                                                                                                                   *tag=
                                                                                                                                                                                                                                                                                                                                                               *tag=
                                                                                                                                                                                                                                                     .20
                                                                                                                                                                                                                                                                                                                                                                                              *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                        *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (ISIS-) ISIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Carras JG, Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 2003-468766/44.
           Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WO2003044163-A2
                                                                                                                                                                                                                                                    modified base
                                                                                                                                                                                                                                                                                                                         modified base
                                                                                                                                                                                                                                                                                                                                                                                   modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        nodified base
                                                                                                                                                                                                                                                                                                                                                      modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                             nodified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            modified base
                                                                                                                                                                                                                        Homo sapiens
                                                                                                                                                   08-SEP-2003
                     20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      responses
                                                                                                                               ACC83613;
  Query Match
          Best Local
Matches 2
                                                                                                   ACC83613/c
                                         ઠ
```

```
Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                   08-SEP-2003
                          Karras JG,
                                                                                       responses.
                                                                                                                                                                                                                                 Query Match
                                                                                                                                                                                                                                                   Matches
                                                                                                                                                                                                                                                                                                                        ACC83598/
à
                                                                                                                                                                                                                                                                                    셤
                                                                                                                                                                                                                                                                                                                                         ö
                       This chimeric phosphorothicate oligonucleotide, having 2-MOE wings and a deoxy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibites 38; inhibition of human Toll-like receptor 4 expression in THP-1 cells. More active oligonucleotides are useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                    Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                                                   /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                 present sequence is that of antisense oligonucleotide ISIS #114645
                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                      Human Toll-like receptor 4 antisense oligonucleotide ISIS #114640.
                                                                                                                                                   ;
                                                                                                                              DB 1; Length 20; 73;
                                                                                                                                                   Indels
                                                                                                                                                    ;
0
                                                                                                                Sequence 20 BP; 5 A; 7 C; 7 G; 1 T; 0 U; 0 Other;
                                                                                                                            0.5%; Scc...
100.0%; Pred. No. ,c,
Example 14; Page 95; 110pp; English
                                                                                                                                                                                                                                                                                                                                                  Location/Qualifiers
                                                                                                                                                                     2304 CCCGGTGTGCCCATTGCTGC 2323
                                                                                                                                                                                                                                                                                                                                                                            /mod_base= OTHER
                                                                                                                                                                                      20 cccediareaccariaciec 1
                                                                                                                                                                                                                                                                                                                                                                                                                                         base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   mod base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  mSc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /*tag= g
/mod_base= m5c
                                                                                                                                                                                                                                  뭠
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               14-NOV-2002; 2002WO-US036390
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                19-NOV-2001; 2001US-00001863
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /*tag= e
/mod_base=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /mod_base=
                                                                                                                                                                                                                                 ACC83585 standard; DNA; 20
                                                                                                                                                                                                                                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                                                      Ø
                                                                                                                                                                                                                                                                                                                                                                                                                                 Д
                                                                                                                                                                                                                                                                                                                                                                                                                                                            υ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ø
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         44
                                                                                                                                           Local Similarity 100.
nes 20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                 *tag=
                                                                                                                                                                                                                                                                                                                                                           1. .20
/*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                          *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         '*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                         /mod
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WO2003044163-A2
                                                                                                                                                                                                                                                                                                                                                           modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                        modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                 modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                nodified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          modified base
                                                                                                                                                                                                                                                                     08-SEP-2003
                                                                                                                                                                                                                                                                                                                                  sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             30-MAY-2003
                                                                                                                                                                                                                                                   ACC83585;
                                                                                                                                  Query Match
                                                                                                                                                    Matches
                                                                                                                                                                                                                                                                                                                                   Ношо
                                                                                                                                                                                                                          ACC83585,
                                                                                                                                                                                                                                   ð
                                                                                                                                                                                      g
```

```
ö
                                                                                                                                                                                                                                                                                                                                    The present sequence is that of antisense oligonucleotide ISIS #114640. This chimeric phosphorothicate oligonucleotide, having 2'-MOE wings and a decoyy ago, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 87% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; ss.
                                                                                                                                                                                expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                       antisense oligonucleotides for modulating Toll-like receptor 4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Human Toll-like receptor 4 antisense oligonucleotide ISIS #114654.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Score 20; DB 1; Length 20;
Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Seguence 20 BP; 7 A; 6 C; 3 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.5%; scc...
100.0%; Pred. No. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2016 ATGCCTGTGCTGAGTTTGAA 2035
                                                                                                                                                                                                                                                                                          Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /*tag= b
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /*tag= c
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /*tag= f
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 20 ATGCCTGTGCTGAGTTTGAA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACC83598 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         mod_base=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ø
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Φ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ъ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1. .20
/*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    *tag=
(ISIS-) ISIS PHARM INC.
                                                  Ξ,
                                                                                                       WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Local Similarity
```

```
Human, Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ACC83587/
    ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                The present sequence is that of antisense oligonucleotide ISIS #114654. This chimeric phosphorothicate oligonucleotide, having 2'-MOE wings and a decay gap, is trargeted to the 3' untranslated region of human Toll-like receptor 4 mRNA. It exhibite 92' inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Th1 or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
                                                                                                                                                                                                                        New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human Toll-like receptor 4 antisense oligonucleotide ISIS #114641
                                                                                                                                                                                                                                                                                                                                                                                                                                                    ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                              Score 20; DB 1; Length 20;
Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                    0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 5 A; 7 C; 4 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                    100.0%; Pred. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      cocation/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        3531 GGGCACTCTTTTAAACGGG 3550
                                                                                                                                                                                                                                                                             Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /mod_base= OTHER
/note= "OTHER = p
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             GGGCACTCTTTAAACGGG 1
            /*tag= g
/mod_base= m5c
                                          /*tag= h
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ACC83586 standard; DNA; 20 BP
                                                                                                                   14-NOV-2002; 2002WO-US036390
                                                                                                                                      19-NOV-2001; 2001US-00001863
                                                                                                                                                                                                                                                                                                                                                                                                                               0.5%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          08-SEP-2003 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                   20; Conservative
                                                                                                                                                          (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                    WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                        Local Similarity
                                                                        WO2003044163-A2
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Key
modified_base
                                modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
                                                                                                                                                                                Karras JG,
                                                                                                                                                                                                                                                           responses.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            20
                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ACC83586
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           셤
```

```
The present sequence is that of antisense oligonucleotide ISIS #114641. This chimeric phosphorothioate oligonucleotide, having 2'-MOB wings and a deoxy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 83% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Human Toll-like receptor 4 antisense oligonucleotide ISIS #114642.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0.5%; Score 20; DB 1; Length 20;
100.0%; Pred. No. 73;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 7 A; 6 C; 5 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2060 CATCATTGGTGTGGGTCC 2079
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Claim 3; Page 95; 110pp; English.
                                                                                                                                                       *tag= d
'mod_base= m5c
/*tag= b
/mod_base= m5c
                                                                                                       mod_base= m5c
                                                                                                                                                                                                                                                               mod_base= m5c
                                                                                                                                                                                                                                                                                                                                          /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                              /*tag= g
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           100.08;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ACC83587 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (first entry)
                                                                                 υ
                                                                                                                                                                                                                                         Ø
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      20; Conservative
                                                                                                                                                                                                                                                                                                                  *tag=
                                                                                                                                                                                                                                      *tag=
                                                                              *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Karras JG, Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WO2003044163-A2
                                                      modified base
                                                                                                                                                                                                           modified base
                                                                                                                                                                                                                                                                                       modified base
                                                                                                                                                                                                                                                                                                                                                                   modified_base
                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               responses,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACC83587;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Local
```

Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.

Location/Qualifiers

modified base

Homo sapiens

Human Toll-like receptor 4 antisense oligonucleotide ISIS #114647

(first entry)

08-SEP-2003

ACC83591 standard; DNA; 20 BP.

```
The present sequence is that of antisense oligonuclectide ISIS #114642. This chimeric phosphorothicate oligonuclectide, having 2'-MOE wings and a decay gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibites 76% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonuclectide is particularly useful for treating or preventing a disease or condition associated with Itall-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                           /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 20 BP; 8 A; 6 C; 5 G; 1 T; 0 U; 0 Other;
                        Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Claim 3; Page 95; 110pp; English.
                                                                  OTHER
                                                                                                                                                                /mod_base= m5c
                                                                                                                                                                                                                                                mod_base= m5c
                                                                                                                                                                                                                                                                                           mSc
                                                                                                                                                                                                                                                                                                                                                                /*tag= g
/mod_base= m5c
                                                                                                                                                                                                          mod_base= m5c
                                                                                                                                                                                                                                                                                                                                    mod_base= m5c
                                                                  /mod_base= OTF
/note= "OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                              14-NOV-2002; 2002WO-US036390
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          19-NOV-2001; 2001US-00001863
                                                                                                                                                                                                                                                                                          mod_base=
                                                                                                                                                                                              υ
                                                                                                                                                                                                                                    ש
                                                                                                                                                                                                                                                                              O
                                                                                                                                                                                                                                                                                                                    44
                                                                                                                                                    *tag=
                                                                                                                                                                                                                                                                             tag=
                                                                                                                                                                                                                                                                                                                       tag=
                                                                                                                                                                                             *tag=
                                                                                                                                                                                                                                    *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                         WO2003044163-A2
                          Key
modified_base
                                                                                                                                                                             modified_base
                                                                                                                                                                                                                                                              modified_base
                                                                                                                                    modified_base
                                                                                                                                                                                                                                                                                                        modified_base
                                                                                                                                                                                                                                                                                                                                                 modified base
                                                                                                                                                                                                                     modified base
Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                   30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                responses.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              The present sequence is that of antisense oligonucleotide ISIS #114647. This chimeric phosphorothioate oligonucleotide, having 2'-MOB wings and a decoxy gap, is targeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 71% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
                                                                                                                                                                                                                          /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central aga region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0.5%; Score 20; DB 1; Length 20; 100.0%; Pred. No. 73; tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 3 A; 4 C; 8 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                                                OTHER
                                                                                                                                                                                                                                                                                                 /*tag= b
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                       /*tag= c
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                        /*tag= e
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                             mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                     1. .20
/*tag= a
/mod_base= (
                                                                                                                                                                                                                                                                                                                                                                               ъ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Local Similarity 100.
                                                                                                                                                                                                                                                                                                                                                                                  *tag≃
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Karras JG, Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                WO2003044163-A2
                                                                                                                                                                                                                                                                                   modified base
                                                                                                                                                                                                                                                                                                                         modified base
                                                                                                                                                                                                                                                                                                                                                                  modified base
                                                                                                                                                                                                                                                                                                                                                                                                         modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
Best Local S:
Matches 20,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   responses
```

2380 AGCACTTCATCCAGAGCCGC 2399

ઠે

; 0

Gaps

; 0

2073 TCGGTCCTCAGTGTGCTTGT 2092

20; Conservative

Matches

Best Local Similarity

20 TCGGTCCTCAGTGTGCTTGT

RESULT 95 ACC83591/c

Н

73;

```
Sequence 20 BP; 4 A; 4 C; 6 G; 6 T; 0 U; 0 Other;
                                                                                                                              ACC83595 standard; DNA; 20
                                                                                                                                                                                                                                  (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        υ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ರ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ø
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WO2003044163-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens
                                                                                                                                                                                                                                  08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Karras JG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       responses
                                                                                                                                                                               ACC83595;
                                                                                                 ACG83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C83595/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C8359/C835
음
```

```
The present sequence is that of antisense oligonucleotide ISIS #114651. This chimeric phosphorothicate oligonucleotide, having 2'-MOE wings and a decay gap, is targeted to the 3' untranslated region of human Toll-like receptor 4 mRNA. It exhibits 91% inhibition of human Toll-like receptor 4 expression in THP-1 cells. It is useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The oligonucleotide is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response, particularly Thl or Th2 responses
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator; phosphorothioate; antisense; 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxymucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e. inflammatory disorders, or conditions involving Th1 or Th2 immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                Human Toll-like receptor 4 antisense oligonucleotide ISIS #114651.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Claim 3; Page 95; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /*tag= a
/mod_base= OTHER
20 AGCACTTCATCCAGAGCCGC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    /*tag= b
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              /mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        19-NOV-2001; 2001US-00001863.
```

0.5%; Score 20; DB 1; Length 20;

Query Match

```
ö
                                                                                                                                                                                    Human; Toll-like receptor 4; receptor; antiinflammatory; immunomodulator;
                                                                                                                                                                                                                                                                            /note= "OTHER = phosphorothioate nucleotides, the oligonucleotide comprises a central gap region of 10 2'-deoxynucleotides, flanked on both sites by 5-nucleotides wings composed of 2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            New antisense oligonucleotides for modulating Toll-like receptor 4 gene expression, particularly useful for preventing, delaying or treating e.g. inflammatory disorders, or conditions involving Thl or Th2 immune
           Gaps
                                                                                                                                                                Human Toll-like receptor 4 antisense oligonucleotide ISIS #114639.
           ..
0
          Indels
           ö
           Mismatches
Best Local Similarity 100.0%; Pred. No. Matches 20; Conservative 0; Mismatch
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Example 14; Page 95; 110pp; English.
                                                                                                                                                                                                                                      Location/Qualifiers
                                                                                                                                                                                                                                                                    'mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                      шSс
                                                                                                                                                                                                                                                                                                                                                                                                                         '*tag= e
'mod_base= m5c
                                                                                                                                                                                                                                                                                                                                           'mod base= m5c
                                                                                                                                                                                                                                                                                                                                                               /*tag= c
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /*tag= g
/mod_base= m5c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /mod_base= m5c
                             2894 GGGCATTTCAACCAACTCAG
                                                  20 GGCATTICAACCAACTCAG
                                                                                                     BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             19-NOV-2001; 2001US-00001863.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          14-NOV-2002; 2002WO-US036390.
                                                                                                                                                                                              phosphorothioate; antisense;
                                                                                                                                                                                                                                                                                                                                                                                           /*tag= d
/mod_base=1
                                                                                          ACC83612/c
ID ACC83612 standard; DNA; 20
                                                                                                                                            (first entry)
                                                                                                                                                                                                                                                             ø
                                                                                                                                                                                                                                                                                                                                                                                                                                                        44
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    *tag≈ h
                                                                                                                                                                                                                                                                                                                                 *tag=
                                                                                                                                                                                                                                                 .20
*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (ISIS-) ISIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                          *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Karras JG, Koller E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2003-468766/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WO2003044163-A2
                                                                                                                                                                                                                                                modified base
                                                                                                                                                                                                                                                                                                                     modified_base
                                                                                                                                                                                                                                                                                                                                                    modified_base
                                                                                                                                                                                                                                                                                                                                                                                  modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          modified base
                                                                                                                                                                                                                                                                                                                                                                                                                 modified_base
                                                                                                                                                                                                                    sapiens
                                                                                                                                            08-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       30-MAY-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              responses.
                                                                                                                       ACC83612;
                                                                                                               à
                                                  g
```

ВР

AAC84782 standard; DNA; 19

```
ó;
                               Ø
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The present invention relates to a novel nucleotide sequence and protein of a new hepatitis E virus HEV-T1 and the application of the nucleotide sequence and protein in diagnosing, preventing and treating hepatitis. The present sequence is a PCR primer described in the exemplification of the invention
        The present sequence is that of antisense oligonucleotide ISIS #114639. This chimeric phosphorothioate oligonucleotide, having 2'-MDE wings and a decoxy gap, is trargeted to the coding region of human Toll-like receptor 4 mRNA. It exhibits 0% inhibition of human Toll-like receptor 4 expression in THP-1 cells. Active oligonucleotides are useful for inhibiting the expression of Toll-like receptor 4 in cells or tissues. The disonate is particularly useful for treating or preventing a disease or condition associated with Toll-like receptor 4, e.g. an inflammatory disorder or a condition involving an immune response,
                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                  .;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Hepatitis E virus; HEV-T1; hepatitis infection; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Score 19.2; DB 1; Length 24; Pred. No. 1.2e+02; 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Hepatitis E virus HEV-T1 sequence related PCR primer #35.
                                                                                                                                                                                                                   Score 20; DB 1; Length 20;
Pred. No. 73;
                                                                                                                                                                                                                                                  0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Hepatitis E virus gene sequence and its application.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 24 BP; 6 A; 4 C; 11 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                   Seguence 20 BP; 1 A; 7 C; 5 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (CHME-) CHINESE MEDICINE & BIOLOGIC PROD APPRAIS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Example 1; Page 15(Disclosure); 34pp; Chinese.
                                                                                                                                                                                                                                100.0%; Prec. ...
tive 0; Mismatches
                                                                                                                                                                                                                                                                                   1943 GATCAAGGACCAGAGCCAGC 1962
                                                                                                                                                                                                                                                                                                               20 garcaagaaccagagcagc
                                                                                                                                                                                                                                                                                                                                                                                               BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ch 0.5%;
1. Similarity 87.5%;
21; Conservative C
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   99CN-00125741.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 99CN-00125741
                                                                                                                                                                                                                      0.5%;
                                                                                                                                                                                                                                                                                                                                                                                             AAI69670 standard; DNA; 24
                                                                                                                                                                                                                                                                                                                                                                                                                                                            (first entry)
                                                                                                                                                                                                                                                     20; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Li H;
                                                                                                                                                                                                                                       Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2001-550442/62
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Zhang H,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   23-DEC-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  23-DEC-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                            10-JAN-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    CN1300771-A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     27-JUN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
Best Local S
                                                                                                                                                                                                                                                                                                                                                                                                                              AA169670;
                                                                                                                                                                                                                        Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Wang Y,
                                                                                                                                                                                                                                                       Matches
                                                                                                                                                                                                                                                                                                                                                                            RESULT
*555555555555
                                                                                                                                                                                                                                                                                     8
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered innate immunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lippolypaccharide) induced chronic airway disease conditions such as systematic inflammatory disease conditions such as systematic inflammatory response syndrome (RNBS), and the respiratory distress syndrome (ARDS), and the respiratory distributions are respiratory distributions are respiratory distributions are respiratory distributions.
                                                                                                                                                                                       TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNJ amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  .;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Score 19; DB 1; Length 19;
Pred. No. 86;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 19 BP; 5 A; 6 C; 3 G; 5 T; 0 U; 0 Other;
                                                                                                                                               Human TLR4 gene exon 4 amplifying forward primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        100.0%; Preq. ...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Schwartz DA, Schutte BC;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   812 CTTTATCCAACCAGGTGCA 830
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1 crrrarccaaccaggraca 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.5%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (IOWA ) UNIV IOWA RES FOUND.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ADB39125 standard; DNA; 19
                                                                                                             (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match 0.5
Best Local Similarity 100.
Matches 19; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                        WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                              Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        10-JUN-1999;
                                                                                                             20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                               21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         receptor 4.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Lorenz E,
                                                                 AAC84782;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (LORE/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 100
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ADB39125/
g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ДX
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ò
```

ö

Gaps

ö

3250 CCATCCTGGTCATTCTCGAACATG 3273 CCATCCTGGCCCTTCTCGACCATG 1

Matches

24

셤 ò

RESULT 99

(first entry)

ss; RT-PCR; reverse transcriptase; primer; myeloid differentiation factor 88; MyD88; atherosclerosis; thrombosis; restenois; angina pectoris; ischaemia; stroke, heart attack; osteonecrosis; colitis; poor kidney function; congestive heart failure; poor blood circulation; slow wound healing; inflammation; infection; claudication; Toll-like receptor 4; TLR-4; human.

Human Toll-like receptor 4 TLR-4 RT-PCR primer #2.

```
angioplasty restenosis; heart disease; PCR; primer; reverse transcription polymerase chain reaction; RT-PCR; human; 88; TLR-4; Toll-like receptor 4.
                                                   vascular disease; Toll-like receptor-4 inhibitor; TLR-4 inhibitor; atherosclerosis; restenosis; inflammation; vasctropic; antiarteriosclerotic; thrombolytic; cardiant; antiinflammatory; antisense therapy; gene therapy; transplant atherosclerosis; vein-graft atherosclerosis; thrombosis; stent restenosis;
                                   Human toll-like receptor (TLR) -4 RT-PCR primer Seg ID4
                                                                                                                                                                                                           24-OCT-2001; 2001US-0335637P.
17-DEC-2001; 2001US-0341359P.
                                                                                                                                                                                          23-APR-2002; 2002US-00128166
                                                                                                                                                      US2003077279-A1
                                                                                                                                     Homo sapiens.
                                                                                                                                                                        24-APR-2003
ADB39125;
```

Arditi M, Rajavashisth T, Shah PK; (CEDA-) CEDARS SINAI MEDICAL CENT.

17-DEC-2001; 2001US-0341359P. 23-APR-2002; 2002US-00128166. 12-DEC-2002; 2002US-00317992.

US2003148986-A1. 17-AUG-2003

```
A system for inhibiting the biological activity of myeloid differentiation factor 88 (MyD88), useful for treating vascular diseases (e.g. atherosclerosis), comprises an intravascular device and MyD88
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          The invention relates to a system for inhibiting the biological activity of myeloid differentiation factor 88 (MyD88) which comprises an intravascular device and a therapeutic composition coated upon the intravascular device, the composition comprising a MyD88 inhibitor. The system or method is useful for treating vascular diseases including etherosclerosis, transplant atherosclerosis, vein-graft atherosclerosis, thrombosis, restenosis, stent restenosis or angioplasty restenosis, thrombosis, restenosis, stent restenosis suffering from angina pectoris, ischaemias, conditions associated with ischaemias including stroke.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              transient ischaemic attacks, heart attack, osteonecrosis, colitis, poor kidney function or congestive heart failure, poor blood circulation to the extremities and complications of poor blood circulation including slow wound healing, inflammation, infections and claudication. The present sequence represents a human Toll-like receptor 4 TLR-4 reverse transcriptase (RT)-PCR primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Example 6; SEQ ID NO 4; 18pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (e.g. atherosclerosis), compris
inhibitor coated on the device.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 2003-897598/82
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ACC74142;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RESULT 102
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ACC74142/
This invention relates to a novel method for the treatment of a vascular disease through the administration of a Toll-like receptor-4 (TLR-4) inhibitor to a mammal. The TLR-4 protein has been linked to several disease such as atherosclerosis, restenosis, inflammation and other. Vascular diseases. Compounds which inhibit the activity of TLR-4, through the inhibition of its receptor, may have vasotropic, the inhibition of its receptor, may have vasotropic, antiatreficaclerotic, thrombollyric, cardiant and antiinflammatory activities. This may also be achieved through antisense therapy or gene therapy. The method or the system of the invention may therefore be conseful for inhibiting or treating a vascular disease, for example atherosclerosis, restenosis, tent restenosis, angioplasty restenosis, or inflammation and other heart disease. The present sequence is that of a constinct which was used for reverse transcription polymerses chain and also the present transcription polymerses chain and also the present sequence is that of a constinct that the activity of the constinct the present sequence is that of a constinct that the constinct the present sequence is that of a constinct that the constinct the constitution of the constitution and the constitution and constitution the constitution and constitution the constitution and constitution the constitution and co
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      thrombosis,
administering
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        PCR primer which was used for reverse transcription polymerase chair reaction amplification of human TLR-4 in the exemplification of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Treating a vascular disease, particularly atherosclerosis, restenosis, stent restenosis or angioplasty restenosis, by a Toll-like receptor-4 (TIR-4) inhibitor to a mammal.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.5%; Score 19; DB 1; Length 19; 100.0%; Pred. No. 86; 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence 19 BP; 5 A; 6 C; 5 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (CEDA-) CEDARS SINAI MEDICAL CENT.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Example 6; Page 10; 21pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      2256 GAAGGGTGCCTCCATTTC 2274
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   GAAGGGGTGCCTCCATTTC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              100.08;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Arditi M, Rajavashisth T,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Best Local Similarity 100.
Matches 19; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2003-615988/58.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADIS3115/c
ID ADIS31
XX ADIS31
XX ADIS31
XX ADIS31
DT 22-APR
```

poor

```
ö
                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                         Mouse, immunomodulator; antibacterial; immunosuppressive; CSF-1; colony stimulating factor-1; septic shock; TLR; toll-like receptor;
                                                                                 ö
                                       Score 19; DB 1; Length 19;
Pred. No. 86;
                                                                                 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                 Reverse primer for detecting murine TLR4 expression.
Sequence 19 BP; 5 A; 6 C; 5 G; 3 T; 0 U; 0 Other;
                                                                                 Mismatches
                                         0.50,
100.0%; Pre-
                                                                                                                        2256 GAAGGGGTGCCTCCATTTC 2274
                                                                                                                                                                                                                                                               ВР
                                                                                                                                             19 GAAGGGGTGCCTCCATTTC
                                                                                                                                                                                                                                                               ACC74142 standard; DNA; 24
                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                            Local Similarity 100.
ses 19; Conservative
                                                                                                                                                                                                                                                                                                                                              11-JUL-2003
```

ADIS3115 standard; DNA; 19

ઠ

22-APR-2004 (first entry)

ADI53115

```
ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                        The invention relates to modulating an immune response in an animal. The method of the invention comprises modulating colony stimulating factor-1 (CSF-1) activity in order to modulate the immune response of the animal. Also disclosed is a pharmaceutical composition comprising a modulator of CSF-1 activity and a pharmaceutical carrier. The method or the pharmaceutical composition is useful for the prophylactic or therapeutic treatment of bacterially-induced septic shock. The sequences given in records ACC74129-ACC74161 represent primers and probes used in an example from the invention to detect murine genes
                                                                                                                                                                                                                    Modulating an immune response in an animal, useful for the prophylactic or therapeutic treatment of bacterially-induced septic shock, by modulating colony stimulating factor-1 (CSF-1) activity in an animal.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   therapeutic agent; endothelial disorder; 3-amino-1; 2-benzisothiazole compound; endothelial disorders; toxaemia; 2-benzisothiazole compound; endothelial disorders; toxaemia; severe toxaemia; toxic shock; haemorrhagic shock; alcock; alcohol induced cirrhosis; adult respiratory distress syndrome; chronic rheumatoid arthritis; ulcerative gastritis; Crohn's disease; glomerulonephritis; infectious carditis; systemic lupus erythematosus; scleroderma; Sjoegren's syndrome; multiple organ failure; autoimmune disease; multiple sclerosis; PCR; primer; ss; human; toll-like receptor 4; TLR4.
                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
 interleukin-12; IL-12; HPRT; hypoxanthine phosphoribosyl transferase;
                                                                                                                                                                                                                                                                                                                                                                                                                                                            ő
                                                                                                                                                                                                                                                                                                                                                                                                                                  0.5%; Score 18.8; DB 1; Length 24; 90.9%; Pred. No. 1.3e+02; tive 0; Mismatches 2; Indel8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human toll-like receptor 4 (TLR4) gene PCR primer #4.
                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 24 BP; 7 A; 4 C; 5 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                           Sester DP;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   625 AACTTAATGTGGCTCACAATCT 646
                                                                                                                                                                                                                                                                      Example, Page 18-19, 58pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AACTCAATGTGGCTCACAATTT 1
                                                                                                                                                                           Sweet MJ, Stacey KJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   31-MAR-2003; 2003WO-JP004108.
                                                                                                      03-OCT-2002; 2002WO-AU001348.
                                                                                                                               03-OCT-2001; 2001AU-00008071.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ADF17210 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                             20; Conservative
                                                                                                                                                    (UYQU ) UNIV QUEENSLAND
                                                                                                                                                                                                  WPI; 2003-381587/36
                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WO2003087072-A1
                                                           WO2003028752-Al.
               PCR; primer; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              12-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             23-OCT-2003
                                                                                   10-APR-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         22
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ADF17210;
                                                                                                                                                                            Hume DA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Matches
                                       Mus
   g
```

```
·;
                                                                                                                                                                                                                                Agents for treating diseases associated with endothelial disorders, toxemia or toll like receptor signaling comprise new or known 3-amino-1,2-benzisothiazole compounds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  endothelial disorders, toxaemia, severe toxaemia, toxic shock, haemorrhagic shock, alcohol induced cirrhosis, adult respiratory distress syndrome, chronic rheumatoid arthritis, ulcerative gastritis, Crohn's disease, glomerulonephritis, infectious carditis, systemic lupus erythematosus, scleroderma, Sjoegren's syndrome, multiple organ failure or autoimmune diseases (e.g. multiple sclerosis). The present DNA sequence represents a PCR primer that was used in the exemplification of
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Constructing a strain of diploid fungal cells in which both alleles of a gene are modified comprises modifying the alleles of a gene in the fungal
                                                                                                                                                                                                                                                                                                                                                                                                                The invention comprises therapeutic agents for preventing or treating diseases associated with endothelial disorders, the agents contain a 3-amino-1,2-benzisothiazole compound. The therapeutic agents of the invention are useful for preventing/treating: diseases associated with
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Diploid fungal cell; allele; gene disruption cassette; promoter replacement fragment; antifungal; fungicide; gene therapy; infection; Candida albicans; identification; primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             m
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      C. albicans specific gene, orf6.2403, identification primer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.5%; Score 18.4; DB 1; Length 20; 95.0%; Pred. No. 1.1e+02; tive 0; Mismatches 1; Indels
                                                                                                                    Mori S;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 20 BP; 4 A; 6 C; 3 G; 7 T; 0 U; 0 Other;
                                                                                                                    Mizuno M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Bussey H;
                                                                                                                                                                                                                                                                                                                                                                 Example 4; SEQ ID NO 7; 180pp; Japanese.
                                                                                                                       Nakamura M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        2784 GCTAAGGGTGAGTAATTCCA 2803
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              20 GCTAAGGGTGAGTAAATCCA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Boone C,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 19-DEC-2003; 2003WO-US040618.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             .9-DEC-2002; 2002US-0434832P
                                                               (MOCH ) MOCHIDA PHARM CO LID.
29-MAR-2002; 2002JP-00132121
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADP98293 standard; DNA; 24
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (ELIT-) ELITRA PHARM INC.
(ELIT-) ELITRA CANADA LTD.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Conservative
                                                                                                                    Satoh T,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Roemer T, Jiang B,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2004-500296/47
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Local Similarity
hes 19; Conserv
                                                                                                                                                                                WPI; 2003-903121/82
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Candida albicans.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WO2004056965-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          23-SEP-2004
                                                                                                                           Purusako S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ADP98293;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 104
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ADP98293/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Matches
```

cells by recombination using a gene disruption cassette and a promoter replacement fragment.

Claim 36; SEQ ID NO 5088; 163pp; English.

The formation labates to a movel method for constructing a sitrain of method comprises and another than the method comprises to a modify and the alleds of a gens side all blood fiftied to method comprises and second laboration using a gene disruption cassette and a gens all blood fiftied to method comprises assembling a collection of diploid fungal cells each of which comprises modified alleds of a gene disruption cassette and the comprises modified alleds of a gene with the same of the comprises of the collection of diploid fungal cells each of which comprises and collection of diploid fungal cells each of which comprises and collection of diploid fungal arrains consortising an independent of a gene disruption cassetter of a manual labes of the collection will be collected to the collection of diploid fungal arrains consortising the diploid strains cuted the collection of diploid fungal arrains consortising method above, and the collection, a nucleic acid molecule, where each micriac aid molecule comprises a muchaer of a diploid fungal arrains consortising the collection of diploid fungal a fundamental and an antifungal agent that in Mystidiable to a target muchocide sequence are modified and are arrains in the collection, a nucleic acid molecule, makes each micriac aid molecule comprises a muchaer of a diploid fungal to all a fungal cell, a purified or isolated muchaer of a diploid fungal to all a fungal cell, a purified or isolated muchaer of a diploid fungal to all and an antifungal agent identifying an antifungal and agent that inhibition of growth or provide an ammalian disease, or correlating changes in the provider of a diploid fungal to all assets of protein of all post of the disease of protein of a diploid fungal to all assets of a d

```
ö
              products as effective targets for therapeutic intervention, and for collecting identified essential genes. This polymuclectide sequence represents an identification primer used in the exemplification of the invention. NOTE: This sequence was downloaded from an electronic sequence
targets for therapeutic intervention, for identifying and validating gene
                                                                                                                                                                                                                                                                                                                                                                                                                         TLR4; toll receptor 4; innate immunity; gram-negative bacteria; sepsis; respiratory distress syndrome; LPS; lipopolysaccharide; asthma; ARDS; chronic airway disease; arthritis; inflammatory disease; SIRS; human; systematic inflammatory response syndrome; pyelonephritis; bronchitis; acute respiratory distress syndrome; gall bladder disease; pneumonia; cystic fibrosis; antibacterial; antiinflammatory; PCR primer; se.
                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Identifying humans at risk of, or having indication associated with altered innate immunity involves detecting or determining whether DNJ amplified from a biological sample encodes a portion of variant toll
                                                                                                                                                                   ö
                                                                                                                                 Similarity 87.0%; Pred. No. 1.5e+02; Observative 0; Mismatches 3; Indels
                                                                                                      Sequence 24 BP; 2 A; 3 C; 7 G; 12 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                Human TLR4 gene exon 4 amplifying forward primer.
                                                                        listing provided on the WIPO website.
                                                                                                                                                                                               488 ATTGACAGGAAACCCCATCCAGA 510
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Lorenz E, Schwartz DA, Schutte BC;
                                                                                                                                                                                                                           23 ATCGACAAGAAACCCATCCAGA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Example 1; Page 31; 97pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               08-JUN-2000; 2000WO-US015723.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             .0-JUN-1999; 99US-00329515.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          IOWA ) UNIV IOWA RES FOUND.
                                                                                                                Query Match
Best Local Similarity 6...
Local 20; Conservative
                                                                                                                                                                                                                                                                                                      AAC84790 standard; DNA; 18
                                                                                                                                                                                                                                                                                                                                                                (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 2001-061872/07.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (LORE/) LORENZ E.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WO200077204-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                  20-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 21-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  receptor 4.
                                                                                                                                                                                                                                                                                                                                    AAC84790;
                                                                                                                                                                                                                                                                         RESULT 105
                                                                                                                                                                                                                                                                                           AAC84790
8888888888
```

The invention relates to human toll receptor 4 (TLR4) nucleic acid and methods to identify polymorphisms at the human TLR4 locus and to identify individuals at risk of, or having, an indication associated with altered inmunity. A variant TLR4 nucleic acid is useful as a diagnostic reagent for detecting a polymorphism in human TLR4 gene. Since the presence of TLR4 mutation is associated with gram-negative sepsis, severity of sepsis, pre-term delivery and respiratory distress syndrome in pre-term infants, agents which alter TLR4 activity are useful for preventing or ameliorating infection by gram-negative bacteria, sepsis induced by gram-negative bacteria, LPS (lipopolysaccharide) induced chronic airway disease, asthma, arthritis, local and systemic inflammatory disease conditions such as systematic inflammatory response syndrome (SIRS) or acute respiratory distress syndrome (ARDS), pyelonephritis, gall bladder disease, pneumonia, bronchitis, chronic

B

```
The present invention relates to a new protein that can be used for avoiding endotoxin shock caused by bacterial infection. The present nucleic acid sequence represents one of a collection (ABKR8338-ABK88345) of synthetic TLR4 PCR primers that were used in the methods of the
                                                                                                                                                                                                                                                                                                                                            An anti-bacterial protein useful for avoiding endotoxin shock caused bacterial infection.
                                                                                                                             Endotoxin shock; bacterial infection; TLR4; PCR; primer; ss.
                                                                                                                                                                                                                                                                                           (NAGO-) ZH NAGOYA SANGYO KAGAKU KENKYUSHO.
                                                                                                                                                                                                                                                                                                                                                                                        Disclosure; Page 6; 16pp; Japanese.
Synthetic TLR4 PCR primer #6
                                                                                                                                                                                                                                       14-DEC-2000; 2000JP-00380561.
                                                                                                                                                                                                                                                                14-DEC-2000; 2000JP-00380561.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WO2003094857-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   19;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               26-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20-NOV-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ADG17647;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 108
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Mus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ð
                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         amplification or hybridisation, e.g. to determine common or disparate ancestry between 2 or more plants, to monitor the genetic contribution of an ancestral plant, to trace the progeny of proprietary plants, in certification of a hybrid plant or to identify the progeny of a back-crossed plant with an ancestral plant
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AAV51401-V51704 are forward PCR primers used to amplify fragments of the Zea mays genome in order to detect polymorphic markers. Such markers can be used in the construction of allele-specific primers and probes for
     obstructive pulmonary disease, local gram-negative bacterial infection and cystic fibrosis. Sequences AAC84776-823 represent PCR primers for amplifying the exons of human TLR4 gene
                                                                                                                                                                                                                                                                                                                                               Polymorphic marker, allele-specific, probe, amplification, PCR primer, hybridisation, plant; hybrid certification; genetic contribution, progeny; back-cross; hybrid; ancestry; corn; ss.
                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     primers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               .
0
                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Brassica species allele-specific oligonucleotide probes and useful for plant breeding.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0.5%; Score 17.4; DB 1; Length 22; 94.7%; Pred. No. 1.6e+02; ive 0; Mismatches 1; Indels
                                                                         Score 18; DB 1; Length 18; Pred. No. 1e+02; 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 22 BP; 10 A; 5 C; 5 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Ä
                                                            Sequence 18 BP; 3 A; 4 C; 3 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sapolsky RJ, Murigneux
                                                                                           100.0%; Pred. ...
                                                                                                                                                                                                                                                                                                                      Zea mays genome forward PCR primer #39
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Example 1; Page 50; 65pp; English
                                                                                                                                          1763 TICATIGGATACGITICC 1780
                                                                                                                                                                     1 rrcarregaracerrrce 18
                                                                                                                                                                                                                                       BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                97WO-US021782
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          96US-0032069P
                                                                                       0.5%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        97US-00813507
                                                                                                                                                                                                                                     AAV51439 standard; DNA; 22
                                                                                                                                                                                                                                                                                           02-FEB-1999 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match 0.5:
Best Local Similarity 94.7
Matches 18; Conservative
                                                                                                  Local Similarity 100.
1es 18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Landry BS,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (AFFY-) AFFYMETRIX INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 1998-333252/29.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  01-DEC-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                            WO9824796-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           02-DEC-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        07-MAR-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       11-JUN-1998.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Lemieux B,
                                                                                                                                                                                                                                                                                                                                                                                                     Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                 Zea mays.
                                                                                                                                                                                                                                                                AAV51439;
                                                                                        Query Match
                                                                                                                  Matches
                                                                                                                                                                                                             RESULT 106
                                                                                                                                                                                                                         ន្តដ្ឋនូន
                                                                                                                                           8
                                                                                                                                                                   용
```

ρλ

```
ô
                                                                                                                                                                                                                                                                                                  differentiation; osteoclast precursor cell; Toll-like receptor ligand; TLR ligand; osteopathic; gene therapy; bone loss; bacterial infection; PCR; primer; ss; RT-PCR; reverse transcription PCR; mouse; murine.
                                                        Gaps
                                                        ö
                       0.5%; Score 17.2; DB 1; Length 22; 86.4%; Pred. No. 1.7e+02; ive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                        Mouse toll-like receptor (TLR) 4 RT-PCR primer SegID9.
Sequence 22 BP; 4 A; 9 C; 1 G; 8 T; 0 U; 0 Other;
                                                                                     2225 GGTAAGGAATGAGCTAGTAAAG 2246
                                                                                                              22 GGTGAGAATGAGCTGGTAAAG 1
                                                                                                                                                                                       ВЪ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           10-MAY-2002; 2002US-0379941P.
14-APR-2003; 2003US-0462859P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                 12-MAY-2003; 2003WO-US014946
                                                                                                                                                                         ADG17647/c
ID ADG17647 standard; DNA; 22
                                                                                                                                                                                                                                               (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (UYPE-) UNIV PENNSYLVANIA.
                                                          Conservative
                                        Local Similarity
```

TTTCCGCTTCCTGGTCTTA 3576

3558 13

ð g RESULT 107

TTTCCGCTTACTGGTCTTA 1

```
This sequence represents an example of an antisense oligonucleotide of the invention. The oligonucleotides are 8-50 nucleotides in length, and are targeted to a nucleic acid encoding human B-raf and which is capable of inhibiting human B-raf expression. The oligonucleotides is used to inhibit the (abnormal) expression of human B-raf, to inhibit hyperproliferation of calls, to treat or prevent an abnormal proliferative condition, e.g. hyperproliferative disorders such as cancer (e.g. of the brain or nervous system), restenosis, psoriasis or a disorder characterised by T-cell activation and growth. They may also be used to diagnose these diseases, as well as atherosclerosis. The oligonucleotides of the invention may be used to distinguish rafeassociated tumours from tumours having other etiologies. The antisense oligonucleotides can also be used to quantify raf expression in assays
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Human; raf; hyperproliferation; neovascularisation; ocular angiogenesis; therapy; cancer; cytostatic; anti-angiogenic; vascular; ophthalmological; antisense; phosphorothioate backbone; B-raf kinase; ss.
                New antisense oligonucleotides - for modulation of human B-raf gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human B-raf kinase antisense oligonucleotide ISIS #14142.
                                                                                                                                                                                                                                                                                                                                                 0.4%; Score 17; DB 1; Length 20;
100.0%; Pred. No. 1.5e+02;
                                                                                                                                                                                                                                                                                                                   Sequence 20 BP; 3 A; 6 C; 1 G; 10 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /note= "Phosphorothioate backbone"
                                                                                                                                                                                                                                                                                                                                                                                0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Location/Qualifiers
                                                              Disclosure; Page 22; 72pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                 1465 TGAAACAAATGAGTGAG 1481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAD44802 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        95WO-US007111.
96US-00756806.
97US-00888982.
98WO-US013961.
                                                                                                                                                                                                                                                                                                                                                                 100.08;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             18-FEB-2000; 2000US-00506073.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        98US-00143214
                                                                                                                                                                                                                                                                                                                                                                                                                                               20 TGAAACAAATGAGTGAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ಹ
                                                                                                                                                                                                                                                                                                                                                                Best Local Similarity 100.
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2002-597918/64.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   nodified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       SISI (-SISI)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         13-DEC-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            31-MAY-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                US6410518-B1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           11-MAY-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      28-AUG-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               25-JUN-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           26-NOV-1996
07-JUL-1997
                                expression.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         06-JUL-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Monia BP;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AAD44802;
                                                                                                                                                                                                                                                                                                                                                  Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Homo
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAD44802/
   셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                  an osteoclast precursor cell comprises contacting the cell with at least one TLR nor the osteoclast precursor cell. This ligand, where the TLR ligand stimulates at least one TLR on the osteoclast precursor cell. The invention may be useful for the development of compounds with an osteopathic activity or ifor gene therapy. The composition and methods are useful in inhibiting differentiation of an osteoclast precursor cell which may be used in the development of therapies to treat patients suffering from bone loss associated with bacterial infection and bone loss resulting from other diseases. These may also be used in identifying a Toll-like receptor that inhibits that of a mouse toll-like receptor (TLR) RT-PCR primer which was used in the exemplification of the invention.
                                                                                                                                                                                      This invention relates to a novel method of inhibiting differentiation of
                                                                          Inhibiting differentiation of an osteoclast precursor cell, useful for treating bone diseases, comprises contacting the cell with a Toll-like receptor (TLR) ligand that stimulates a TLR on the osteoclast precursor
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Antisense oligonucleotide; B-raf; human; inhibitor; T-cell activation;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 hyperproliferative disorder; cancer; restenosis; psoriasis; atherosclerosis; raf-associated tumour; diagnosis; therapy; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.5%; Score 17.2; DB 1; Length 22;
86.4%; Pred. No. 1.7e+02;
iive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human B-raf kinase antisense oligonucleotide Isis#14142.
                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence 22 BP; 4 A; 9 C; 1 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /*tag= a
/note= "phosphorothioate bases"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          2225 GGTAAGGAATGAGCTAGTAAAG 2246
                                                                                                                                                       Example; SEQ ID NO 9; 72pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            22 GGTGAGAAATGAGCTGGTAAAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       AAX21943 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       97US-00888982
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       98WO-US013961
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    18-MAY-1999 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 . .20
/*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      PHARM INC
                                            WPI; 2003-903937/82.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 1999-120502/10.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Local Similarity
tes 19; Conser
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      SISI (-SISI)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WO9902167-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         06-JUL-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       07-JUL-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           21-JAN-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Monia BP;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AAX21943
                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match
                Choi Y;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Matches
ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 셤
```

Gaps

ö

0; Indels

```
capable of
                                                                      The present invention relates to novel antisense oligonucleotides which are targetted to nucleic acids encoding human raf proteins and capable of inhibiting raf expression. The invention also relates to methods of inhibiting hyperproliferation of cells which involves contacting the hyperproliferation of cells which involves contacting the hyperproliferation of the invention. The method is useful for treating cancer, angiogenesis or neovascularisation, especially ocular angiogenesis or neovascularisation. The present DNA sequence is an antisense oligonucleotide targetted to human B-raf kinase
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Preventing and/or treating conditions associated with raf expression, such as hyperproliferative disorders, atherosclerosis and tumors, using
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             tumour metastasis; human; raf; raf expression inhibitor; cytostatic; antiarteriosclerotic; antisense-therapy; hyperproliferative disorder; atherosclerosis; tumour; b-raf kinase; antisense oligonucleotide; ss.
                                                                                                                                                                                                                                                                                       Gaps
Treating cancer, angiogenesis or neovascularization by administering antisense oligonucleotides targeted to human raf sequences.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         such as hyperproliferative disorders, atherosclerosis and tumors,
antisense oligonucleotide modulation of human raf gene expression.
                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                        0.4%; Score 17; DB 1; Length 20;
00.0%; Pred. No. 1.5e+02;
                                                                                                                                                                                                                                                                                   0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human b-raf kinase antisense oligonucleotide seq id 65.
                                                                                                                                                                                                                          Sequence 20 BP; 3 A; 6 C; 1 G; 10 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                      0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ID NO 92; 41pp; English
                                             Example 18; Col 26; 41pp; English
                                                                                                                                                                                                                                                                                                                    TGAAACAAATGAGTGAG 1481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           94US-00250856.
95WO-US007111.
96US-0756806.
97US-00888982.
98WO-US013961.
98US-00143214.
                                                                                                                                                                                                                                                                                                                                                                                                                            BP.
                                                                                                                                                                                                                                                                        100.08;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 14-JUN-2002; 2002US-00173225
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     2002US-00057550
                                                                                                                                                                                                                                                                                                                                               rgaaacaaargagrgag
                                                                                                                                                                                                                                                                                                                                                                                                                            ADF09796 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (first entry)
                                                                                                                                                                                                                                                                                       17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2003-863446/80.
                                                                                                                                                                                                                                                                         Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (MONI/) MONIA B
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        US2003119769-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             31-MAY-1995;
26-NOV-1996;
07-JUL-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      25-JAN-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               31-MAY-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    12-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       28-AUG-1998;
18-FEB-2000;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    26-JUN-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          06-JUL-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Monia BP;
                                                                                                                                                                                                                                                                                                                    1465
                                                                                                                                                                                                                                                                                                                                                  20
                                                                                                                                                                                                                                                                                                                                                                                                                                                      ADF09796;
                                                                                                                                                                                                                                                                                      Matches
                                                                                                                                                                                                                                                                                                                                                                                            RESULT 11.
ADF09796/
                                                                                                                                                                                                                                                                                                                                                                                                                            Š
                                                                                                                                                                                                                                                                                                                                               유
```

The invention describes a method of preventing or treating tumour metastasis in an animal comprising administering to the animal an oligonucleotide 8-50 nucleotides in length, which is targeted to encoding human raf and capable of inhibiting raf expression. Also

ö

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                which is targeted to mRNA encoding human c-raf, a-raf or b-raf (raf is a protein kinase playing a regulatory role in signal transduction, regulating cell proliferation and has been implicated in lung carcinoma), and which is capable of inhibiting raf expression. Also included is a composition comprising the oligonucleotide and a pharmaceutically acceptable carrier. The antisense oligonucleotide is useful for inhibiting the expression of human raf in human cells or tissues by contacting the human cells or tissues with the oligo. The oligo. is also is useful for treating or preventing a disease or condition associated
disclosed are raf oligonucleotides, nucleic acids, proteins and compositions used in the methods of the invention. The oligonucleotides have cytostatic and antiarteriosclerotic properties, are useful as Rafinhibitors and in antisense-therapy. The methods and compositions of the present invention are useful for preventing and/or treating conditions associated with raf expression, such as hyperproliferative disorders, atherosclerosis and tumours. This sequence represents a human b-raf
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Novel antisense oligonucleotide which is targeted to mRNA encoding human raf and which is capable of inhibiting raf expression, useful for treating or preventing hyperproliferative conditions such as cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   The invention relates to an oligonucleotide 8-50 nucleotides in length
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human; ss; antisense; c-raf; a-raf; b-raf; protein kinase; cancer; signal transduction; cell proliferation; lung carcinoma; cytostatic; antisense gene therapy; chemotherapeutic agent; angiogenesis; hyperproliferative condition; neovascularisation; ocular angiogenesis.
                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Antisense oligonucleotide targeting human b-raf, ISIS14142.
                                                                                                                                                                                                     ch 0.4%; Score 17; DB 1; Length 20; 1 Similarity 100.0%; Pred. No. 1.5e+02; 17; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                      Sequence 20 BP; 3 A; 6 C; 1 G; 10 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Example 18; Page 14; 42pp; English.
                                                                                                                                     kinase antisense oligonucleotide.
                                                                                                                                                                                                                                                                               1465 TGAAACAAATGAGTGAG 1481
                                                                                                                                                                                                                                                                                                                                                                                                             ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    95WO-US007111.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                2002US-00057550
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   94US-00250856
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    97US-00888982
98WO-US013961
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       98US-00143214
2000US-00506073
                                                                                                                                                                                                                                                                                                                  20 reaaacaaareacreac
                                                                                                                                                                                                                                                                                                                                                                                                             ACD42185 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2003-503332/47
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Δ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         US2003032607-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ш
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (MONI/) MONIA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         28-AUG-1998;
18-FEB-2000;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                25-JAN-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      05-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      31-MAY-1995;
26-NOV-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              13-FEB-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        07-JUL-1997
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           06-JUL-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Monia BP;
                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACD42185;
                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                               Local
                                                                                                                                                                                                                                                                                                                                                                             RESULT 112
                                                                                                                                                                                                                                               Matches
                                                                                                                                                                                                                                                                                                                                                                                              ACD42185,
                                                                                                                                                                                                                                                                                                                                                                                                                 8888888888
```

Gaps

; 0

Length 20;

```
Bergeron MG, Picard FJ, Ouellette M,
                                                                                                                                                                                                                                                  (IDII-) IDI INFECTIO DIAGNOSTIC INC
                                                                                                                                                                                                                                                                                                      Claim 21; Page 93; 167pp; English
                                                                                                  1465 TGAAACAAATGAGTGAG 1481
                                                                                                                                                                                                                                                                                           antibiotic resistance gene(s).
                                                                                                                                      AAV37060 standard; DNA; 20 BP
                                                                                                            20 TGAAACAAATGAGTGAG 4
                                                                                                                                                                                                                              97WO-CA000829.
                                                                                                                                                                                                                                        96US-00743637
                                                                                                                                                           04-SEP-1998 (first entry)
                                                                                        17; Conservative
                                                                                                                                                                                                                                                                       WPI; 1998-286967/25.
                                                                                  Local Similarity
                                                         human raf mRNA
                                                                                                                                                                                                                             04-NOV-1997;
                                                                                                                                                                                                         WO9820157-A2
                                                                                                                                                                                                                    14-MAY-1998
                                                                                                                                                                                               Synthetic
                                                                                                                                                AAV37060;
                                                                              Query Match
                                                                                                                                 Matches
                                                                                                                            RESULT 113
8888888888888888888
                                                                                                           g
                                                                                                   ઠ
```

```
method of the invention comprises modulating colony stimulating factor-I (CSP-1) activity in order to modulate the immune response of the animal. Also disclosed is a pharmaceutical composition comprising a modulator of CSF-1 activity and a pharmaceutical carrier. The method or the pharmaceutical carrier. The method or the pharmaceutical for the prophylactic or therapeutic readment of bacterially-induced septic shock. The sequences given in records ACC74129-ACC74161 represent primers and probes used in an example from the invention to detect murine genes
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   The invention relates to modulating an immune response in an animal. The
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Modulating an immune response in an animal, useful for the prophylactic or therapeutic treatment of bacterially-induced septic shock, by modulating colony stimulating factor-1 (CSF-1) activity in an animal.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Mouse, immunomodulator; antibacterial; immunosuppressive; CSF-1; colony stimulating factor-1; septic shock; TLR; toll-like receptor; interleukin-12; IL-12; HPRT; hypoxanthine phosphoribosyl transferase;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match 0.4%; Score 16.8; DB 1; Length 20; Best Local Similarity 90.0%; Pred. No. 1.6e+02; Matches 18; Conservative 0; Mismatches 2; Indels
                                                                                                              0.4%; Score 16.8; DB 1; Length 2
90.0%; Pred. No. 1.6e+02;
tive 0; Mismatches 2; Indels
     determine the bacterial resistance to antibiotics
                                                          Sequence 20 BP; 4 A; 3 C; 7 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 20 BP; 4 A; 2 C; 9 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Forward primer TLR4 for semi-quantitative PCR.
                                                                                                                                                                                                                            2378 CCAGCACTTCATCCAGAGCC 2397
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    548 ACAGAAGCTGGTGGCTGTGG 567
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1 AGAGAATCTGGTGGCTGTGG 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Stacey KJ,
                                                                                                                                                                                                                                                                                 20 ccaccacricarcascacre 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Example; Page 19; 58pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                            ACC74158 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               03-OCT-2002; 2002WO-AU001348.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   03-OCT-2001; 2001AU-00008071
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (first entry)
                                                                                                  Ouery Match
Best Local Similarity 90.0°
Matches 18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (UYOU ) UNIV QUEENSLAND.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Hume DA, Sweet MJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2003-381587/36
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WO2003028752-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     PCR; primer; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        11-JUL-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        10-APR-2003.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ACC74158;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Mus sp.
                                                                                                                                                                                                                                                                                                                                                                  RESULT 114
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 115
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADD93829
     ន្តដូល
                                                                                                                                                                                                                                                                                                                                                                                                                                                      ò
                                                                                                                                                                                                                                                                                 셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     g
                    chemotherapeutic agent to a human or cells of the human, where the expression of raf is abnormal expression, and the condition is a hyperproliferative condition such as cancer, angiogenesis or neovascularisation (preferably ocular angiogenesis or neovascularisation). The oligo, is also useful for inhibiting hyperproliferation of cells. The oligos, are also useful as tools, for example for detecting and determing the role of raf expression in various cell functions and physiological processes and conditions and for diagnosing conditions associated with raf expression and for purposes. The present sequence is an antisense oligonucleotide targeting
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           PCR primers AAV37060-61 and AAV37062-63 are used to amplify antibiotic resistance gene aac6'-IIa. They are used in the course of the invention. The specification describes the use of probes and/or amplification primers which are specific, ubiquitous and sensitive for determining presence and amount of nucleic acids from a bacterial antibiotic resistance gene and specific bacterial and fungal species in any sample suspected of containing the bacterial or fungal nucleic acids, where each of the nucleic acid or variant or part comprises a selected target region bybridisable with the probes or primers. The method of use comprises contacting the sample with the probes or primers and detecting the presence of hybridised probes or amplified products as an indication of the presence of the specific bacterial or fungal species and bacterial antibiotic resistance genes. The methods and products can be used to detect and identify the bacterial and fungal species and genera and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gaps
expression of raf by administering it in combination with
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            identification and quantification of bacteria, fungi and bacterial
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Detection; bacterial antibiotic resistance gene; bacteria; fungal species; identification; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Use of oligo:nucleotide primers and probes - for detection,
                                                                                                                                                                                                                                                                                                                                                                                                                         Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     PCR primer for antibiotic resistance gene aac6'-IIa.
                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 3 A; 6 C; 1 G; 10 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                    0.4%; Score 17; DB 1; Le
100.0%; Pred. No. 1.5e+02;
iive 0; Mismatches 0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Roy PH;
```

Sester DP;

ö

Gaps

ö

```
Mus
```

```
apoptosis-associated tyrosine kinase, apoptosis, nerve cell differentiation, nerve cell proliferation; nerve cell differentiation, nerve cell proliferation; nerve cell division; drug screening; transgenic animal; apoptosis disorder; nerve regeneration; neurological disorder; hepatitis; hepatotropic; anti-inflammatory; virucide; cytostatic; reverse transcription-PCR; RT-PCR; primer; ss.
                                                                                                                                                                                                                                                                                          Mouse; murine; AF06972; mAF06972; HJ06972; KIAA1079; AATYK homologue;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              New polynucleotides encoding proteins useful for treating disorders associated with cell growth, division and death and hepatitis.
                                                                                                                                                                                                                      Mouse AF06972 (mAF06972) RT-PCR primer, SEQ ID NO:18.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 5 A; 3 C; 5 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Nishimura S;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                primers used in an example of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Example 7; SEQ ID NO 18; 142pp; Japanese.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Satoh S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (KAZU-) KAZUSA DNA RES INST.
(FUJI ) FUJISAWA PHARM CO LTD.
ADD93829 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               26-MAR-2003; 2003WO-JP003713.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        26-MAR-2002; 2002JP-00086843
                                                                                                                                                 29-JAN-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Miyoshi S, Zenkoh J,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 2003-865124/80
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WO2003080836-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              02-OCT-2003.
                                                                            ADD93829
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     gb.
```

```
The invention relates to a splice variant of human HJ06972 (KIRA1079), designated AF06972 (ADD93815), and nucleic acids encoding it (ADD93814). Gengared to the HJ06972 (DAD (ADD93812), AF06972 (DAD contains an additional 94 bp exon (ADD93816) between bases 4623-4624 of HJ06972 and its splice variant AF06972 are related to apoptosis-associated tyrosine kinase (AATYK), a protein which is involved in apoptosis and the liferentiation or proleiferation of nerve cells, and are likely to have similar activity. The invention also relates to vectors and host cells comprising AF06972 DNA sequences, the recombinant production of AF06972 polypeptides, antibodies specific for AF06972 in immunoassay method using the antibodies, a method for detecting apoptosis regulatory activity, a method for evaluating a compound for its ability to induce activity, a method for evaluating a compound for its ability to induce activity, a method for evaluating a compound for its ability to induce activity, and related compounds and methods of the invention may be used in the treatment of disorders associated with apoptosis, nerve cell growth and division, and hepatitis. Sequences ADD93830 nerve cell growth and division, and hepatitis. Sequences ADD93830 nerve
```

```
ö
                                      Gaps
                                   ;
0
0.4%; Score 16.8; DB 1; Length 20; 90.0%; Pred. No. 1.6e+02; tive 0; Mismatches 2; Indels
                                        18; Conservative
        Query Match
Best Local Similarity
Matches 18; Conserval
```

ö

2093 AGTATCTGTTGTAGCAGTTC 2112 ||||| |||||||||| |||||| AGTACCTGTTGTAGAAGTTC 20

RESULT 116 ABZ87635/c

RESULT 117 ABD23865/c

```
The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation codon, coding region, 5, or 3, end genomic flanking regions, 5 and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiasthmatic, hypotensive, case in antisense gene therapy. The composition may have a immunosuppressive, and cytostatic activity. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in a subject of the composition of depleting or for enhancing the prophylactic or therapeutic respiratory effect of an antisinflammatory steroid in a subject, for reducing or depleting levels of ... receptor, producing bronchodilation, increasing levels of adenosine receptor, producing bronchodilation, increasing levels of ubiquinone or lung surfactant in a subject is tissue, or treating bronchoconscription, lung allergies, or a respiratory disease or condition. Note: The sequence data for this patent is not represented in the printed specification, but was obtained in electronic format directly from WIPO
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or
                                                                                                                                                                          Human, antisense, lung dysfunction, nasal airway dysfunction, antinflammatory steroid; ubiquinone; antiinflammatory, antiallergic; antiatlematic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Aguilar
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0.4%; Score 16.8; DB 1; Length 20; 90.0%; Pred. No. 1.6e+02; ive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Pabalan J,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Sequence 20 BP; 7 A; 3 C; 5 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               at ftp.wipo.int/pub/published_pct_sequences
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Disclosure; SEQ ID NO 2877; 872pp; English.
                                                                                                                                                                                                                                                                                                 inflammation; respiratory disease; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Katz E,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1857 CAGCATTTTCCAAGTAGTCT 1876
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Nyce JW, Li Y, Sandrasagra A, K
Miller S, Tang L, Shahabuddin S;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20 CAGCATTTTCCAAGTACTGT 1
                                                                                                                                    Human oligonucleotide sequence.
BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           24-APR-2001; 2001US-0286137P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              23-APR-2002; 2002WO-US013135.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (EPIG-) EPIGENESIS PHARM INC.
ABZ87635 standard; DNA; 20
                                                                                       (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match 0.4
Best Local Similarity 90.0
Matches 18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2003-229219/22.
                                                                                                                                                                                                                                                                                                                                                                                         WO200285308-A2.
                                                                                                                                                                                                                                                                                                                                              Homo sapiens.
                                                                                          17-OCT-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                      31-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ubiquinone
                                             ABZ87635;
                                                                                                                                                                                                                                                                                                   lung
  g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ò
```

ABD23865 standard; DNA; 20

(first entry) 29-JUL-2004 Human myosin X-derived oligonucleotide SEQ ID 2877.

Human, antisense, bronchoconstriction, allergy, hyposecretion, pain, respiratory tract inflammation, adenosine sensitivity, lung, cancer, surfactant depletion, antiallergic, antiinflammatory; antiasthmatic, analgesic, hypotensive, immunosuppressive, cytostatic, cystic fibrosis; beta-adrenergic agonist; respiratory disease, pulmonary vasoconstriction, respiratory distress syndrome; allergic rhinitis; pulmonary hypertension, emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; 88; primer.

Homo sapiens.

WO200285309-A2

31-OCT-2002

23-APR-2002; 2002WO-US013143.

24-APR-2001; 2001US-0286036P.

(EPIG-) EPIGENESIS PHARM INC

Aguilar D; Pabalan J, Katz E, Li Y, Sandrasagra A, Ka Tang L, Shahabuddin S; Miller S, Nyce JW,

WPI; 2003-093058/08.

S Pharmaceutical composition for treating asthma, has antisense oligonucleotide containing less percentage of adenosine, targeted trucleic acids associated with lung airway or lung dysfunction, and bronchodilating agent.

Claim 15; SEQ ID NO 2877; 763pp; English.

This invention describes a novel composition (a) a first active agent, comprising oligonuclectides, effective for alleviating bronchoconstriction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung dysfunction or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonuclocities, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antiinflammatory, antiasthmatic, pulmonary obstruction, and/or bronchoconstriction and/or lung inflammation, allergies and/or surfactant hypoproduction are associated with a disease or condition such as pulmonary vasoconstriction, inflammation, allergies, asthma, impeded respiration, respiratory distress syndrome, pain, oystic fibrosis, allergic rhinitis, pulmonary hypertension, emphysema, chronic obstructive pulmonary disease, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of the oligonucleotides into products that free adenosine into the system the oligonucleotides into products that free adenosine into the system e.g., lung, brain, heart, kidney, etc, tissue environment and thereby, to prevent any unwanted effects due to it analgesic, hypotensive, immunosuppressive and cytostatic activity, is a beta-adrenergic agonist. The composition is useful for preventing or reating a respiratory, lung or malignant disease. The administered composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypeptide present in the lungs. The

ö

Gaps

ö

0.4%; Score 16.8; DB 1; Length 20; 10.0%; Pred. No. 1.6e+02; ve 0; Mismatches 2; Indels

90.06;

Local Similarity

Query Match

Conservative

18,

Matches

1323 TTCAAAGGTTGCTGTTCTCA 1342

||||||| ||||||||||| TTCAAATGTTGCTGCTGTTCTGA

셤

ઠ

ADH66115 standard; DNA; 20 BP

RESULT 119

ADH661.

ADH66115

20

Sequence 20 BP; 4 A; 3 C; 4 G; 9 T; 0 U; 0 Other;

Sequence 20 BP; 7 A; 3 C; 5 G; 5 T; 0 U; 0 Other;

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             The invention comprises an antisense oligonucleotides that are targeted antisense acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
                                                                                                                                                                                                                                                                                                                                       antisense oligonucleotide; glucocorticoid receptor; infection; infilammation; tumour formation; diabetes; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2-methoxyethyl; 2-MOB.
                                                                                                                                                                                                                                                                                                       Human glucocorticoid receptor-specific antisense oligonucleotide #3302
                                   Gaps
                                   ö
 Length 20;
0.4%; Score 16.8; DB 1; Length 2 90.0%; Pred. No. 1.6e+02; ive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Claim 4; SEQ ID NO 3302; 985pp; English
                                                                    1857 CAGCATTTTCCAAGTAGTCT 1876
                                                                                                        20 CAGCATTTCCAAGTACTGT
                                                                                                                                                                                               ВЪ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20-MAY-2002; 2002US-0381857P.
                                                                                                                                                                                               ADH66468 standard; DNA; 20
                                                                                                                                                                                                                                                                     (first entry)
                                     Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Crosby SD, Nalseth AE;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2004-035034/03.
 Query Match
Best Local Similarity
Matches 18; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WO2003099215-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20-MAY-2003;
                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens.
                                                                                                                                                                                                                                                                     25-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        04-DEC-2003
                                                                                                                                                                                                                                  ADH66468;
                                                                                                                                                            RESULT 118
                                                                                                                                                                                ADH66468
                                                                                                                                                                                                              8
```

```
The invention comprises an antisense oligonucleotides that are targeted antisense acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                             New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 antisense oligonucleotide; glucocorticoid receptor; infection; inflammation; tumour formation; diabetees; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2-methoxyethyl; 2'-MOE.
                                                                                  antisense oligonucleotide; glucocorticoid receptor; infection; inflammation; tumour formation; diabetees; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2'-methoxyethyl; 2'-MOE.
                                                  Human glucocorticoid receptor-specific antisense oligonucleotide #2949
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human glucocorticoid receptor-specific antisense oligonucleotide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     0.4%; Score 16.8; DB 1; Length 20; 90.0%; Pred. No. 1.6e+02; tive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Seguence 20 BP; 5 A; 3 C; 4 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Claim 4; SEQ ID NO 2949; 985pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1324 TCAAAGGTTGCTGTTCTCAA 1343
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               rcaaargrigcrgricrgaa 20
                                                                                                                                                                                                                                                                                                                   20-MAY-2002; 2002US-0381857P.
                                                                                                                                                                                                                                                                                  20-MAY-2003; 2003WO-US016084.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ADH67804 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              25-MAR-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                          Crosby SD, Nalseth AE;
                                                                                                                                                                                                                                                                                                                                                         (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                              WPI; 2004-035034/03.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Best Local Similarity
                                                                                                                                                                                                           WO2003099215-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WO2003099215-A2
                                                                                                                                                                         Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens
                                                                                                                                                                                                                                              04-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                04-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ADH67804;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     RESULT 120
ADH67804/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Matches
셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ò
```

ö

Gaps

```
ö
                                                                                                                                                                                                                                                                                The invention comprises an antisense oligonucleotides that are targeted to nucleic acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothioate backbone.
                                                                                                                                                                             New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Polymorphism, human, inhibitor; cancer; treatment; cell growth; LOH; cell viability; loss of heterozygosity; precancerous condition; ASI; allele specific inhibitor; somatic cell; diagnosis; prevention; atherosclerotic plaque; premalignant metaplastic lesion; andometriosis; dysplastic lesion; benign tumnour; polycystic kidney disease; transplant; graft versus host disease; malignant cell removal; bone marrow; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Identifying target genes for allele-specific drugs - used for diagnosis, prevention and treatment of, e.g. cancers, atherosclerotic plaque,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 16.8; DB 1; Length 20; 90.0%; Pred. No. 1.6e+02; ive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 20 BP; 12 A; 0 C; 1 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                     Claim 4; SEQ ID NO 4638; 985pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Stanton VP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  3297 CTGATATATTTTTTA 3316
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    20 Chaarararrrrararrra 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        BP,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        97US-0041057P.
              20-MAY-2003; 2003WO-US016084
                                              20-MAY-2002; 2002US-0381857P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       98WO-US005419
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Human polymorphic region 59.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAZ25870 standard; DNA; 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Local Similarity 90.0
Les 18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          VARI-) VARIAGENICS INC
                                                                                                                Crosby SD, Nalseth AE,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Ledley FD,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 1998-521232/44.
                                                                                                                                                 WPI; 2004-035034/03
                                                                              (PHAA ) PHARMACIA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WO9841648-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       19-MAR-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20-MAR-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          30-NOV-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     24-SEP-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Housman D,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AAZ25870;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 121
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAZ25870
d
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ð
```

dysplastic lesions, endometriosis or graft versus host disease. 2314 CCATTGCTGCCAACATCATC 2333 Example 14; Fig 1; 605pp; English. 1 CCATTGCTGCCAACATGACC 20 AAV80034 standard; DNA; 20 BP 98WO-EP002593. 97GB-00008851. 16-MAR-1999 (first entry) Best Local Similarity 90.0 Matches 18; Conservative (GENZ ) GENZYME UK LTD. Homo sapiens WO9849324-A2 30-APR-1998; 30-APR-1997; 27-JAN-1998; 05-NOV-1998. AAV80034; Synthetic Query Match ઠ

This invention describes a novel method for identifying an inhibitor potentially useful for treatment of cancer, where the inhibitor is active on a gene vital for cell growth or viability, and where the gene is subject to loss of heteroxygosity (LOH) in a cancer. The inhibitor is used for preventing the development of cancer in a patient having a precancerous condition, by administering to the patient a first allele specific inhibitor (ASI) targeted to an allele of a first essential gene present in cells of the precancerous condition, where the normal somatic cells of the precancerous for the first gene, the inhibitor is active on at least one but less than all allelic forms of the gene present in a population and targets only one allelic form present in the concern and an allele of the precancerous conditions of the gene concern, at the diagnosis, premailignant metaplastic or dysplastic cancers, atherosclerotic plaques, premailignant metaplastic or dysplastic or dissing tumours, endometriosis, polycyclic kindry disease, and AAZ25812-Z26825 represent graft versus host disease. The method can also be used to remove malignant cells from bone marrow transplants. AAZ25812-Z26825 rej human polymorphic sites described in the method of the invention

Sequence 21 BP; 5 A; 8 C; 3 G; 5 T; 0 U; 0 Other;

Gaps ö 0.4%; Score 16.8; DB 1; Length 21; 90.0%; Pred. No. 1.7e+02; ative 0; Mismatches 2; Indels

ö

Primer intU2 for SSCP analysis of PMM2 exon 5A.

Phosphomannomutase-2; PMM2; CDG1; mutation; human; transgenic; assay; carbohydrate-deficient glycoprotein syndrome type 1; drug screening; Jaeken disease; single-strand confirmation polymorphism; SSCP; prenatal diagnosis; PCR primer; ss.

Matthijs G;

WPI; 1999-024063/02.

New DNA encoding human phosphomannomutase or its fragments - used to detect mutations associated with carbohydrate-deficient glycoprotein syndrome-1, particularly for prenatal diagnosis.

the coding region of the human helicase-moi gene. The antisense oligomucleotides of the invention are useful for inhibiting the spression of human helicase-moi in cells or tissues, and for treating a helicase-moi-associated condition. The antisense oligomucleotides of the invention may also be used to delay infection, inflammation and tumour

The invention comprises antisense oligonucleotides which are targeted

Claim 3; Col 43-44; 52pp; English.

Novel antisense compound for modulating expression of human helicase-moi and for treating inflammation, specifically hybridizes to a specific region in nucleic acid molecule encoding the human helicase-moi.

WPI; 2002-749291/81.

Claim 5; Page 64; 104pp; English.

the nucleotide sequence encoding the protein. The DNA or its fragments are used to detect mutation in the PMMZ genes that are associated with the carbohydrate-deficient glycoprotein syndrome type I (CDGI). The sequences can also be used to detect expression of PMMZ-related CDNA; to express PMMZ or its mutants; and to create transgenic animals for use in drug screening and for studying expression pathways. The expressed proteins are used to screen for agents that modulate activity of PMMZ, for therapy and to raise specific antibodies (for detecting PMMZ or its mutants, in competitive or capture assays). Biochemical assays for phophomannomutaes activity are used to identify possible carriers of CDGI (Jaeken disease). Measuring enzymatic activity in foetal cells (or in parental leucocytes if such cells are unavailable) and detecting mutations in the PMMZ gene makes possible a better prenatal diagnosis of CDGI. Sequences AAV80026-41 represent primers used in PCR and singlestand confirmation polymorphism (SSCP) analysis of the 8 exons of PMMZ gene. These primers are used to determine the SSCP mutations in the PMMZ ö invention relates to a human phosphomannomutase-2 (PMM2) protein and Gaps Human; antisense gene therapy; phosphorothioate backbone; antisense oligonucleotide; helicase-moi gene; inflammation; ss; helicase-moi-associated condition; infection; tumour formation; 2-MOE nucleotide; 2'-methoxyethyl nucleotide. ö 0.4%; Score 16.4; DB 1; Length 20; 94.4%; Pred. No. 1.8e+02; ve 0; Mismatches 1; Indels Human helicase-moi inhibiting oligonucleotide #1. Seguence 20 BP; 8 A; 4 C; 5 G; 3 T; 0 U; 0 Other; 1640 CACAGAGCTGAGAACTT 1657 BP. 10-MAY-2001; 2001US-00853768. 10-MAY-2001; 2001US-00853768. 94.48; ABT13876 standard; DNA; 20 13-FEB-2003 (first entry) Matches 17; Conservative (ISIS-) ISIS PHARM INC Local Similarity Ward DT, Watt AT; US6444466-B1. Homo sapiens 3-SEP-2002. ABT13876; Query Match ABT13876/ 셤 à

Gaps

ö

Indels Length

Score 16.4; DB 1; Pred. No. 1.8e+02; 0; Mismatches 1;

0.48;

94.48;

```
interference. The present sequence represents a human helicase-moi antisense oligonucleotide.
                                                               Sequence 20 BP; 4 A; 4 C; 3 G; 9 T; 0 U; 0 Other;
                                                                                                                                                                                             613 AAACTTTGAAAGAACTTA 630
                                                                                                                              Best Local Similarity 94.4
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                            Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Local
                                                                                                                                                                                                                                                                                                 RESULT 125
ADH65626
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Best Loc
Matches
      ន្តដូល
                                                                                                                                                                                                                                                                                                                                            ద
                                                                                                                                                                                             ò
                                                                                                                                                                                                                                  셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        à
                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   The invention relates to antisense oligonucleotides, compositions and methods for modulating the expression of helicase-moi. The oligonucleotides are used in treating an animal having a disease or condition associated with helicase-moi, such as inflammation, a hyperproliferative disorder or a condition that arises from RNA-mediated
  formation. The present DNA sequence represents a human helicase-moi gene antisense oligonuclectide of the invention. NOTE: The present DNA sequence has a phosphorothioate backbone, bases 1-5 and 16-20 are 2'-methoxyethyl (2'-MOE) nucleotides
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ,
'
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        New antisense oligonucleotides for modulating helicase-moi expression, useful for diagnosing, preventing or treating diseases or conditions associated with helicase-moi, e.g. inflammation or hyperproliferative disorders.
                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Cytostatic; Antisense therapy; ss; human; helicase-moi; inflammation; hyperproliferative disorder; RNA-mediated interference; probe.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /mod_base= Other
/note= "Phosphorothioate linkages. All cytidines are
methylcytidines"
                                                                                                                                                                                             ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  note= "2'-methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 nucleotides
                                                                                                                                                   Length 20;
                                                                                                                                                                                             1; Indels
                                                                                                                                               Score 16.4; DB 1;
Pred. No. 1.8e+02;
0; Mismatches 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /*tag= a
/mod_base= Other
/note= "2'-methoxyethyl (2'-MOE)
                                                                                                          Seguence 20 BP; 4 A; 4 C; 3 G; 9 T; 0 U; 0 Other,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Human helicase-moi, antisense oligonucleotide #1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Claim 3; SEQ ID NO 14; 56pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /*tag= c
/mod_base= Other
                                                                                                                                                                                                                                    613 AAACTTTGAAAGAACTTA 630
                                                                                                                                                                                                                                                                           N
                                                                                                                                                                                                                                                                                                                                                                                      BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         31-MAY-2002; 2002US-00160632
                                                                                                                                                 Query Match

Best Local Similarity 94.4%;
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   10-MAY-2001; 2001US-00853768
                                                                                                                                                                                                                                                                               AAACTCTGAAAGAACTTA
                                                                                                                                                                                                                                                                                                                                                                                 ADN60081 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Д
                                                                                                                                                                                                                                                                                                                                                                                                                                                                    01-JUL-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 16. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ...20
*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 2003-898586/82.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Watt AT;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (WARD/) WARD D T. (WATT/) WATT A T.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             US2003176380-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     18-SEP-2003
                                                                                                                                                                                                                                                                           19
                                                                                                                                                                                                                                                                                                                                                                                                                           ADN60081;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Ward DT,
                                                                                                                                                                                                                                                                                                                                            RESULT 124
                                                                                                                                                                                                                                                                                                                                                              ADN60081/6
XX
AC ADN66
XX
AC ADN66
XX
ADN60
XX
A
    8888888
                                                                                                                                                                                                                                                                           g
                                                                                                                                                                                                                                      ð
```

```
;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   The invention comprises an antisense oligonucleotides that are targeted to nucleic acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-NOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
                                                                                                                                                                                                                                                                                                                                                                                                                                  antisense oligonucleotide; glucocorticoid receptor; infection; inflammation; tumour formation; diabetes; obesity; tumour formation; thyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2 - methoxyethyl; 2'-MOE.
                                                                                                                                                                                                                                                                                                                                                                             #2460
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                             Human glucocorticoid receptor-specific antisense oligonucleotide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             .;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Score 16.4; DB 1; Length 20;
Pred. No. 1.8e+02;
0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 3 A; 5 C; 3 G; 9 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Claim 4; SEQ ID NO 2460; 985pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1340
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20
                   N
                                                                                                                                                                                      BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20-MAY-2003; 2003WO-US016084.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20-MAY-2002; 2002US-0381857P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1 Similarity 94.4%;
17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1323 TTCAAAGGTTGCTGTTCT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        rrcaaargrigcrerrcr
19 AAACTCTGAAAGAACTTA
                                                                                                                                                                                      ADH65626 standard; DNA; 20
                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Crosby SD, Nalseth AE;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 2004-035034/03.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO2003099215-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens
                                                                                                                                                                                                                                                                                                              25-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               04-DEC-2003.
                                                                                                                                                                                                                                                  ADH65626;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           m
```

RESULT 126

```
New antisense oligonuclectides for modulating never in mitosis, gene a (NIMA)-related kinase 6 expression, useful for diagnosing, preventing or treating diseases associated with the kinase, e.g. hyperproliferative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The present invention relates to antisense compounds targeted to a nucleic acid encoding human never in mitosis gene a-related kinase 6 (MIMA-related kinase 6). The antisense compound comprises an antisense oligomuclectide that specifically hybridises with the nucleic acid and inhibits the expression of NIMA-related kinase 6. The antisense oligomuclectide is a chimeric oligomuclectide. The antisense oligomuclectide comprises at least one modified internucleoside linkage, preferably a phosphorochioate linkage. It also comprises at least one modified sugar moiety, preferably a 2'-0-methoxyethyl (2'-MOE) sugar moiety. The antisense oligomucleotide further comprises at least one modified nucleobase, preferably a 5-methylcycosine. The antisense oligomucleotide further comprises at least one modified nucleobase, preferably a 5-methylcycosine. The antisense oligomucleotide further comprises
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         hyperproliferative disorders, e.g. cancer. The present sequence represents a human NIMA-related kinase 6 DNA target sequence for an
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0.4%; Score 16.4; DB 1; Length 20;
94.4%; Pred. No. 1.8e+02;
Ive 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 20 BP; 5 A; 7 C; 4 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Example 15; SEQ ID NO 122; 51pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1439 ACATCTGGATTTCCAGCA 1456
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1 ACATCTGGATGTCCAGCA 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             B
                                                                                                       16-NOV-2002; 2002US-00295471.
                                                                                                                                                           16-NOV-2002; 2002US-00295471.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Local Similarity 94.4%;
nes 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADO55815 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ಥ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               antisense oligonucleotide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1. .20
/*tag=
                                                                                                                                                                                                            (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                WPI; 2004-389184/36.
US2004097441-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 12-AUG-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens
                                                      20-MAY-2004.
                                                                                                                                                                                                                                                                  Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ADO55815;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AD055815,
     ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          The invention comprises an antisense oligonucleotides that are targeted antisense acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
                                                                                                                                                                                                                               antisense oligonucleotide; glucocorticoid receptor; infection; inflammation; tumour formation; diabetees; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2 - methoxyethyl; 2 - MOE.
                                                                                                                                                                                Human glucocorticoid receptor-specific antisense oligonucleotide #4092.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Antisense therapy; human; NIWA-related kinase 6;
never in mitosis gene a-related kinase 6; hyperproliferative disorder;
cancer; cytostatic; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%; Score 16.4; DB 1; Length 20;
44.4%; Pred. No. 1.8e+02;
ve 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human NIMA-related kinase 6 DNA target sequence #30.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 3 A; 4 C; 4 G; 9 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Claim 4; SEQ ID NO 4092; 985pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1323 TTCAAAGGTTGCTGTTCT 1340
                        ADH67258 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ADO55876 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20-MAY-2003; 2003WO-US016084
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20-MAY-2002; 2002US-0381857P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               2 rrcaaargrigcrigricr
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               94.48;
                                                                                                                             25-MAR-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   12-AUG-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (PHAA ) PHARMACIA CORP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Crosby SD, Nalseth AE;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2004-035034/03.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                   WO2003099215-A2
                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      04-DEC-2003
                                                                            ADH67258;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
Best Local 8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADO55876
```

ö

Gaps

ö

```
/note= "This oligonuclectide has a phosphorothioate backbone and 2'-methyoxyethyl (2'-MOS) wings at the 5' and 3' ends, which are 5 muclectides in length at each end. All cytidine residues are 5-methylcytidines"
                                                                                                     Antisense therapy; human; NIMA_related kinase 6;
never in mitosis gene a-related kinase 6; hyperproliferative disorder;
cancer; cytostatic; phosphorothioate; ss.
                                                                               Human NIMA-related kinase 6 DNA, antisense oligonucleotide #38
                                                                                                                                                                                            Location/Qualifiers
```

.

Homo sapiens

RESULT 127

ठ 셤 AD055876

```
The present invention relates to antisense compounds targeted to a nucleic acid encoding human never in mitosis gene a-related kinase 6 (NIMA-related kinase 6). The antisense compound comprises an antisense oligonuclectide that specifically hybridises with the nucleic acid and inhibits the expression of NIMA-related kinase 6. The antisense coligonuclectide is a chimeric oligonuclectide. The antisense oligonuclectide comprises at least one modified internucleoside linkage, preferably a phosphorothicate linkage. It also comprises at least one moditied sugar moderty, preferably a 2.0-methoxyethyl (2.-MOS) sugar moderty, preferably a 5.-methylcytosine. The antisense conjectide further comprises at least one modified nucleobase, preferably a 5-methylcytosine. The antisense conjectide further comprises such as hyperproliferative disorders, e.g. cancer. The present sequence represents an antisense oligonucleotide used in the examples of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      murine; PCR; ss; antidiabetic; mouse; beta-cell hypofunction; diabetes; type II diabetes; non-insulin dependent diabetes mellitus; NIDDM; primer.
                                                                                                                                                                                                                      New antisense oligonucleotides for modulating never in mitosis, gene a (NIMA) -related kinase 6 expression, useful for diagnosing, preventing or treating diseases associated with the kinase, e.g. hyperproliferative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           PCR primer used to amplify a murine gene SeqID 18.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence 20 BP; 4 A; 4 C; 7 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  y Match
Local Similarity 94.4%; Pred. No. 1.8e+02;
hes 17; Conservative 0; Mismatrhen
                                                                                                                                                                                                                                                                                                       Example 15; SEQ ID NO 52; 51pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1439 ACATCTGGATTTCCAGCA 1456
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ٣
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       18-OCT-2002; 2002JP-00304524.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          17-OCT-2003; 2003WO-JP013311.
                                                            16-NOV-2002; 2002US-00295471.
                                                                                            16-NOV-2002; 2002US-00295471.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 20 ACATCTGGATGTCCAGCA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADN14299 standard; DNA; 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (first entry)
                                                                                                                           (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (SANY ) SANKYO CO LTD
                                                                                                                                                                                           WPI; 2004-389184/36.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WO2004035830-A1.
US2004097441-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Mus musculus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            15-JUL-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          29-APR-2004
                                20-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ADN14299;
                                                                                                                                                                                                                                                                            disorders
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                                                                                                                                                                          treating
                                                                                                                                                           Dobie KW
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Best Loc?
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 129
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              8
```

```
this invention relates to a novel method for evaluating substances for their effectiveness in treating beca-cell hypofunction. Specifically, it refers to the beta cells of the pancreas islets that normally function to synthesise and secrete insulin in response to glucose in the blood. The present invention describes measuring the expression levels of at least one of eight specific genes in the presence of a test substance, to indicate its effectiveness in treating beta-cell hypofunction.
Accordingly, this method can be used in the treatment of diabetes, in particular type II diabetes (or non-insulin dependent diabetes, in wIDDM). This oligonuclectide is a murine PCR primer given in an exemplification of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Polymerase chain reaction; PCR; primer; amplify; human; T cell receptor; beta chain; TCR; myelin basic protein; BP; autoantigen; encephalitogen; experimental autoimmune encephalomyelitis; BAR; multiple sclerosis; MS; autoimmune disease; neurological disease; cerebrospinal fluid; therapy; central nervous system; complementarity determining region; CDR; T lymphocyte; optical nerve damage; anterior chamber inflammation; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Methods for diagnosis and immune-related therapy of autoimmune diseases partic. multiple sclerosis, by detecting marker T cell receptor V gene bias and treating patients with selected V beta peptide(s).
                                                                Evaluating substances for treating functional failure in B-cells caused by diabetes.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AAT10575 and AAT10576 represent amplification primers specific for the human T cell receptor beta (TCRbeta) chain of the myelin basic protein
                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                           .;
0
                                                                                                                                                                                                                                                                                                                                                                                       ch 0.4%; Score 16.4; DB 1; Length 21; 1 Similarity 94.4%; Pred. No. 1.9e+02; 17; Conservative 0. Minnach.
                                                                                                                                                                                                                                                                                                                                                                      Sequence 21 BP; 7 A; 8 C; 2 G; 4 T; 0 U; 0 Other;
Fujiwara T;
                                                                                                                          Example 4; SEQ ID NO 18; 142pp; Japanese
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Vandenbark AA, Offner H, Buenafe A;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (CONN-) CONNECTIVE THERAPEUTICS INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Example 1; Page 26; 62pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                GITCTACATCAAATGCCC 770
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Cbeta-specific primer H3Cbeta3,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 cricracarcaaarccc 20
   Oshima K,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    94US-00270634.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 95WO-US008086
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AAT10575 standard; DNA; 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WPI; 1996-087679/09.
   Taguchi T,
                                      WPI; 2004-365167/34
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WO9601329-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   26-JUN-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      01-JUL-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               18-JAN-1996.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              05-SEP-1996
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAT10575;
                                                                                                                                                                                                                                                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    753
                                                                                                                                                                                                                                                                                                                                                                                                                                  Local
   Koga T,
                                                                                                                                                                                                                                                                                                                                                                                                                                  Best Loca
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT 130
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAT10575
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              g
```

```
encephalomyelitis (EAE), and is the leading candidate as an encephalicygen involved in multiple scalenosis. (MS). By detecting the presence of a marker TCR variable (W) gene bias in a body fluid which encapsulates all or a portion of the target organ, an autoimmune disease (such as a neurological disease) in a human can be identified. This method can also be carried out to detect the presence of a biased motification of the pathogenic antigen in a nontarget tissue or organ. By analysing the Wosta gene repertorie of crebrospinal fluid (CP), and determining the Wosta gene repertorie of crebrospinal fluid (CP), and determining the presence of a Wosta gene bias, an immune-related disease that targets the central nervous system can be disease or organ of the disease, and selection is carried out by identifying a Wosta gene bias in a body fluid that is not the target tissue or organ of the disease, and selecting an immunogenic peptide to the women entarity determining region 2 (CDR2) of a V gene peptide on the surface of a T lymphocyte in the CF of a patient and administering a peptide corresponding to the Wosta gene bias. Ms can be treated by identifying the surface of a T lymphocyte in the CF of a patient and administering a peptide corresponding to this region. These methods can also be used for the diagnosis and immune-related therapy of optical nerve damage and an analyse of a contractor chamber inflammation as well as other human neurological
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                diseases
             %X323232323232323232323233
```

Sequence 21 BP; 4 A; 5 C; 7 G; 5 T; 0 U; 0 Other;

```
Gaps
                                 ö
0.4%; Score 16.2; DB 1; Length 21; 85.7%; Pred. No. 2e+02; ative 0; Mismatches 3; Indels
                                                                2496 CTGCTCAGGCAGCAGGTGGAG 2516
                                                                                             21
                                 18; Conservative
                   Best Local Similarity
   Query Match
                                 Matches
                                                                8
```

ö

1 CTGCTCAGGCAGTATCTGGAG AAV08680 standard; DNA; 21 BP 15-FEB-1999 (first entry) AAV08680; 셤

Primer ATP/17FB for human ACE gene.

PCR primer; human; ACE; angiotensin converting enzyme; angiotensinogen; cardiovascular status; ACT; ATI; type 1 angiotensin II receptor; stroke; polymorphic pattern; blood pressure; electrocardiographic profile; cardiac condition diagnosis; myocardial infarction; atherosclerosis; hypertension; cardiovascular disease; ss.

Homo sapiens. Synthetic

WO9845477-A2

15-0CT-1998

98WO-IB000475 01-APR-1998; 97US-0042930P. 04-APR-1997;

(EURO-) EURONA MEDICAL AB

WPI; 1998-568361/48.

Norberg LT, Andersson MK, Lindstroem PHR;

The invention relates to a method of detecting Kawasaki disease by detecting an increase in Vbeta6.5 or Vbeta6.5/Vbeta2.1 positive T-cells. The sequences AAA95531-A95626 represent primers and probes used to PCR amplify and detect the level of expression of Valpha and Vbeta genes in T

Example 1; Page 6; 36pp; Japanese.

Score 16.2; DB 1; Length 21; Pred. No. 2e+02;

0.4%;

Query Match Best Local Similarity

,

Sequence 21 BP; 8 A; 6 C; 4 G; 3 T; 0 U; 0 Other;

cells in Kawasaki disease

Assessing cardiovascular status in humans by polymorphic analysis - of genes for angiotensin converting enzyme, angiotensinogen and angiotensin II receptor, used to diagnose predisposition to disease and to predict effect of therapy.

Example 1; Page 32; 71pp; English.

```
This sequence represents a PCR primer for the human ACE (angiotensin converting enzyme) gene, and can be used in the method of the invention. The method is for assessing cardiovascular status in humans by determining the sequence of at least one polymorphic site in the ACE determining the sequence of at least one polymorphic site in the ACE angiotensin converting enzyme), AGT (angiotensinnogen) and/or ATI (type 1 angiotensin converting enzyme), AGT (angiotensinnogen) and/or ATI (type 1 angiotensin II receptor) genes, and comparing the polymorphic pattern with that in patients with predetermined markers of status. The method is used to assess blood pressure or electrocardiographic profile, to diagnose a cardiac condition such as (allent) myocardial infarction (MI), bypertension, atherosclerosis or stroke. They can also be used to predict antagonists, diuretics, alpha- or beta-adrenergic receptor antagonists, etc. It is also used to identify susceptibility to cardiovascular can income the genes are used to screen for cardiovascular agents. The nucleic acids contained in the library can be is used as source of probes
                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Detection of Kawasaki disease factor, useful for the diagnosis of
Kawasaki disease, comprises detecting an increase in Vbeta6.5 positive T-
                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Detection; diagnostic; Kawasaki disease; T-cell; PCR primer; probe;
                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                                                    0.4%; Score 16.2; DB 1; Length 21;
85.7%; Pred. No. 2e+02;
tive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                      Sequence 21 BP; 4 A; 8 C; 2 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                TCR Valpha 2 subfamily probe VA02-2.
                                                                                                                                                                                                                                                                                                                                                                                                                         27 AGTGAGGATGATGCCAGGATG 47
                                                                                                                                                                                                                                                                                                                                                                                                                                              21 AGTGAGGTTGATGCCAGAAAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      98JP-00341661.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      98JP-00341661.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAA95550 standard; DNA; 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (SHIO ) SHIONOGI & CO LTD
                                                                                                                                                                                                                                                                                                                                                                                         18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     gene expression; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2000-477722/42.
                                                                                                                                                                                                                                                                                                                                                                       Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     JP2000157297-A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      01-DEC-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       01-DEC-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               31-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      13-JUN-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AAA95550;
                                                                                                                                                                                                                                                                                                                                                        Query Match
                                                                                                                                                                                                                                                                                                                                                                         Best Local
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          cells.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RESULT 132
AAA95550/c
                                                                                                                                                                                                                                                                                                                                                                                      Matches
셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ઠ
```

10001863-3.sl.rng

```
The invention relates to a novel method of assessing the cardiovascular status in an individual and to newly identified polymorphisms in the genes encoding angiotensin-converting enzyme (ACE), angiotensin II receptor type 2 (ATI), angiotensin II receptor type A and beta-adrenergic receptors 1 and 2. The method comprises determining the sequence at one or more polymorphisms from the individual with a reference polymorphic positions within these genes, and comparing the pattern of polymorphisms from the individual with a reference polymorphic pattern ob polymorphisms from the individual with a reference polymorphic pattern ob polymorphic may predict a popularity of a individual exhibiting a predetermined cardiovascular disease status. The polymorphic markers are useful for determining the predisposition of an individual to cardiovascular disease status of an individual to receive and stroke. They are also useful for predicting the likely cardiovascular status of aptient given a predicting the likely cardiovascular status of aptient given a treatment regimen comprising administration of cardiovascular drugs of the genes comprising administration of a treatment regimen. Fragments of the genes comprising a polymorphic site may be used as provides a basis for predicting the outcome of a treatment regimen. Fragments of the genes comprising a polymorphic site may be used as primars and probes for detecting genetic polymorphic pattern reduces or they encode are useful in the screening. The genes, and the proteins change. Determination of an individual's polymorphic pattern reduces or eliminate patients from clinical trials who are predicted to be nonceptual at a risk for an adverse response, to a particular response, or at a risk for an adverse response, to a particular
ö
                                                                                                                                                                                                                                                                                                                                                                     polymorphism; polymorphic marker; cardiovascular disease;
myocardial infarction; unstable angina; hypertension; atherosclerosis;
stroke; prognosis; drug screening; treatment outcome; human; PCR primer;
Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Assessing cardiovascular status in humans involves comparing test polymorphic pattern comprising polymorphic positions within genes encoding specific proteins, with reference polymorphic pattern.
;
0
                                                                                                                                                                                                                                                                                                                                                Angiotensin II receptor type 1 gene; AT1; regulatory region;
Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Jonsson L;
                                                                                                                                                                                                                                                                                                         Human AT1 regulatory region PCR primer, SEQ ID NO:108
3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Lindstrom PHR,
Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Example 1; Page 53; 126pp; English.
                                        429
                                                                             Н
ö
                                        409 TGCTGGATTTATCCAGGTGTG
                                                                             21 recreearriarceaccrers
                                                                                                                                                                                 AAA38308 standard; DNA; 21 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          98US-0104286P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    99WO-IB001678
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Norberg LT, Andersson MK,
                                                                                                                                                                                                                                                                     21-AUG-2000 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (EURO-) EURONA MEDICAL AB.
18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2000-318010/27.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO200022166-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      13-OCT-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          14-OCT-1998;
14-OCT-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20-APR-2000.
                                                                                                                                                                                                                            AAA38308;
Matches
                                                                                                                                                              g
                                        8
```

```
ö
identify polymorphic patterns so that the adverse results can be correlated with a sub-population of the test population, permitting exclusion of such sub-populations from the treatment group. Beneficial drugs can be approved for use in the appropriate population, thereby decreasing the number of patients required for a clinical trial, which in turn decreases the duration and cost of such trials. Sequences AAA3826-A38315 represent PCR primers used in an exemplification of the invention to amplify short fragments of the human angiotensin II receptor type I (AT1) gene regulatory region (AAA38331) for sequence determination
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Assessing disease status in individual by determining sequence(s) at one
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          or more polymorphic positions within the human genes encoding the protein(s) involved in physiological pathway associated with treatment
                                                                                                                                                                                                                                                                                                                                                                                                                                         Human ACE, AGT and AT1 genes polymorphisms PCR primer SEQ ID NO: 108.
                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human, genetic polymorphism, disease diagnosis, treatment, cancer, cardiovascular system, nervous system, glaucoma; PCR primer, ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sanders R;
                                                                                                                                                                                                                        ;
0
                                                                                                                                                                                          0.4%; Score 16.2; DB 1; Length 21; 85.7%; Pred. No. 2e+02; ive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Olaisson E,
                                                                                                                                                             Sequence 21 BP; 4 A; 8 C; 2 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Jonsson L,
                                                                                                                                                                                                                                                    47
                                                                                                                                                                                                                                                                               AGTGAGGTTGATGCCAGAAG 1
                                                                                                                                                                                                                                                    27 AGTGAGGATGATGCCAGGATG
                                                                                                                                                                                                                                                                                                                                                        ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               99WO-IB000497.
99US-0126243P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     23-MAR-2000; 2000WO-GB001102.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 99US-0126046P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           99US-00471890
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Lindstrom PHR, Norberg LT,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (GEMI-) GEMINI GENOMICS AB.
                                                                                                                                                                                                                                                                                                                                                        AAC61308 standard; DNA; 21
                                                                                                                                                                                                                                                                                                                                                                                                                 (first entry)
                                                                                                                                                                                                       1 Similarity 85.7
18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2000-638268/61.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WO200056922-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                23-MAR-1999;
24-MAR-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                 30-JAN-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 23-MAR-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             23-DEC-1999;
                                                                                                                                                                                                                                                                                                                                                                                    AAC61308;
                                                                                                                                                                                              Query Match
Best Local S
                                                                                                                                                                                                                                                                                                                              RESULT 134
                                                                                                                                                                                                                           Matches
                                                                                                                                                                                                                                                                                                                                          AAC61308,
       88888888888888
                                                                                                                                                                                                                                                                                 g
                                                                                                                                                                                                                                                                                                                                                                       ò
```

The present invention is related to methods for determining the polymorphic pattern of an individual and using the results to determine their risk of a number of diseases, including cancer, cardiovascular diseases, glaucoma and nervous system disorders such as depression and neurodegenerative diseases. In addition, the methods can be used to determine the effects of different types of treatment for individuals, and thus enables appropriate therapies to be prescribed. The PCR primers shown in sequences AAC61201-C61371 were all used to demonstrate the

Example 1; Page 61; 141pp; English.

Sequence 21 BP; 4 A; 8 C; 2 G; 7 T; 0 U; 0 Other;

the invention

10001863-3.81.rng

ABA05093 standard; DNA; 21 BP

RESULT 136 ABA05093

```
The present sequence is one of a number of primers used in a method for detecting a mutation or a polymorphism in the human ATM gene, which is associated with the disease ataxia telangiectasia, or a polyexonic eukaryotic gene of at least 4 kb. The method uses an improved version of single-stranded conformation polymorphism (SSCP) electrophoresis that allows electrophoresis of two or three amplified segments in a single lane. The method is useful for screening large, complex polyexonic eukaryotic genes such as the ATM gene for mutations and polymorphisms. The new mutations and polymorphisms in the ATM gene are useful for performing more accurate screening of human DNA samples for mutations, for distinguishing mutations from polymorphisms, and for improving the efficiency of automated screening method: The meethod provides a screening method of genes for multiple polymorphisms and mutations at enemy points and cukaryotic genes, having mutations and polymorphisms at many points and not merely at one or a few hot spote. Note: the SEQ ID assigned to this sequence in the disclosure and claims of the the specification is one
                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Detecting a mutation or polymorphism in human ataxia telangiectasia gene or polyexonic eukaryotic gene, involves using mega-single stranded comformation polymorphism analysis.
                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                  Human, ATM, ataxia telangiectasia; mutation detection;
single-stranded conformation polymorphism; SSCP; electrophoresis;
                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.4%; Score 16.2; DB 1; Length 21;
85.7%; Pred. No. 2e+02;
iive 0; Mismatches 3; Indels
0.4%; Score 16.2; DB 1; Length 21;
85.7%; Pred. No. 2e+02;
iive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       number lower than the number given in the sequence listing
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 21 BP; 13 A; 2 C; 5 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                            Human ATM gene exon 31 reverse primer.
                                                                           27 AGTGAGGATGATGCCAGGATG 47
                                                                                                         21 AGTGAGGTTGATGCCAGAAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Claim 7; Page 53; 118pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            21-JUL-2000; 2000WO-US020011.
                                                                                                                                                                                                         AAF60173 standard; DNA; 21
                                                                                                                                                                                                                                                                                      (first entry)
                                       18; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (REGC ) UNIV CALIFORNIA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WPI; 2001-168574/17.
                 Best Local Similarity
Matches 18; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO200107660-A1
                                                                                                                                                                                                                                                                                                                                                                                                      PCR primer; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   23-JUL-1999;
                                                                                                                                                                                                                                                                                      27-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        01-FEB-2001,
                                                                                                                                                                                                                                                 AAF60173;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gatti RA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match
 Query Match
                                                                                                                                                                                           AAF60173
                                                                                                             셤
                                                                                                                                                                                                               ò
```

```
The present invention provides the protein and coding sequences of human dipeptide aminopeptidase 28. The sequences can be used in the treatment of cancer, nosobaemia, HIV infection, immunological diseases and inflammation. The present sequence is a coding sequence fragment
                                                                                                                                                                                                                                                        /partial
/note= "the sequence contains no start or stop codon"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             nervous system; Thce gene; CofE protein; tubulin-specific chaperone;
                                                                                                          Human, dipeptide aminopeptidase 28; cancer; nosohaemia; cytostatic;
anti-HIV; immunosuppressive; antiinflammatory; HIV infection;
immunological disease; inflammation; gene therapy; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                New polypeptide for treating malignant tumors and HIV infection, comprises the human dipeptide aminopeptidase.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match 0.4%; Score 16.2; DB 1; Length 21; Best Local Similarity 85.7%; Pred. No. 2e+02; Matches 18; Conservative 0; Mismatches 3; Indels
                                                                                Dipeptide aminopeptidase 28 DNA related oligonucleotide #8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 described in the exemplification of the invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 21 BP; 9 A; 1 C; 5 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Murine pmn sequence tagged site PCR primer #2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Disclosure; Page 5(Disclosure); 11pp; Chinese.
                                                                                                                                                                                                                                                                                                                                                                                                                 (JINP-) JINPENG BIO TECH CO LTD SHANGHAI.
                                                                                                                                                                                                                           /*tag= a
/product= "AAM47329"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1127 AAGGGTAAAAGACTTTTCTTA 1147
                                                                                                                                                                                                 Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1 AAGGGTAAAAGACGTTATTTA 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               BP.
                                                                                                                                                                                                                                                                                                                                                          03-FEB-2000; 2000CN-00112673.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ADK00166 standard; DNA; 21
                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                           Tong
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2002-034896/05.
P-PSDB; AAM47329.
                                                                                                                                                                                                                                                                                                                                                                                                                                             Wang J, Wang S,
                                                                                                                                                                                                                                                                                                                                                                                    03-FEB-2000;
                                                                                                                                                                    Unidentified.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        03-JUN-2004
                                                     22-FEB-2002
                                                                                                                                                                                                                                                                                                   CN1307129-A.
                                                                                                                                                                                                                                                                                                                               08-AUG-2001.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADK00166;
                          ABA05093;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     RESULT 137
셤
```

ö

ö

Gaps

ö

1842 AAAAAACAGGAACTACAGCAT 1862

ઠે

Conservative

Local Similarity nes 18; Conserv

Best Loca Matches

AAAAAACAGGAAGAACAGGAT 21

short interfering nucleic acid; siNA; downregulation; inhibition; SCD; stearcyl-CoA desaturase; RNA interference; anorectic; antidiabetic; antiatreiosclerotic; cytostatic; virucide; obesity; diabetes; atherosclerosis; cancer; viral infection; drug screening; genetic engineering; pharmacogenomic; gene mapping; ss.

Stearoyl-CoA desaturase siNA oligonucleotide SEQ ID NO:11.

29-JAN-2004 (first entry)

```
alpha-tubulin; beta-tubulin; progressive motor neuropathy; pmn;
chromosome 13; neuroprotective; nootropic; antiparkinsonian;
degenerative nerve disease; nerve injury; intoxication;
Alzheimer's disease; barkinson's disease; neuropathy; multiple sclerosis;
                                                                                                                                                                                                                                          This invention describes a novel isolated nucleic acid used in a test
                                                                                                                                                                                                       therapeutic agents for degenerative nerve diseases
                                                                                                                                           (MEDL-) MEDLANOVA GES MEDIZINISCHE INNOVATIONEN.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  342 GACCTGAGCTTTAATCCCCTG 362
                                                                                                                                                                                              Thee gene,
                                                                                                                                                                                                                         Example 1; Fig 1; 34pp; German.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   GACCTGAGTTTGAATCCCCAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ADE27067 standard; RNA; 19 BP
                                                                                                        31-JUL-2003; 2003WO-DE002589
                                                                                                                         26-AUG-2002; 2002DE-01039961
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Conservative
                                                                                                                                                                                               mutant form of the
                                                                                                                                                                             WPI; 2004-227242/21.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Local Similarity
es 18; Conser
                                                                     WO2004020661-A2.
                                    ss; primer; PCR
                                                                                       11-MAR-2004
                                                                                                                                                            Sendtner M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ADE27067
                                                    Mus sp.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 138
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ADE27067
ID ADE2
XX
AC ADE2
원
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ð
```

```
This interaction describes a novel isolated nuclear acid used in a rest of system. The nucleic acid is a mutant form of the murine Thee gene which encodes the Octs protein, a tubulin-specific chaperone essential for complexation of alpha- and beta-tubulins. The mutant gene has a T to G alteration at position 1682, resulting in Gly rather than Trp at C- terminal position 1682, resulting in Gly rather than Trp at C- terminal position 1682, resulting in Gly rather than Trp at C- contexted with the identified diagnostic agent and binding determined, contexted with the identified diagnostic agent and binding determined, qualitatively or (semi) quantitatively. The mutant Thee gene may be diagnosed by in vitro hybridisation to mutation-specific probes or by contexted with test compound and either the survival rate or the contexted with test compound and either the survival rate or the formation of embryonal tubulin isoforms, class III beta-tubulin, ordered tubulin aggregates, microtubuli and/or axons determined and optionally compared with results of controls and/or of cells treated with as compound. Compounds are selected if they increase cell survival or reference compound. Compounds are selected if they increase cell survival or or modify formation of tubulin and/or axons. The novel mutation was identified in punypmn mice by typing a 31 cM region of chromosome 13. The products of the invention have neuroprotective, nootropic and antiparkinsonian activity and can be used and its derived peptide to identify agents for diagnosis of degenerative nerve diseases and nerve diseases but mutation and multiple sclerosis. These or is made are also useful in diagnosing and treating the disorders. This sequence represents an inverse PCR.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   and treating the disorders. This sequence represents an inverse PCR primer used to amplify murine pmn candidate region sequence tagged sites (STS) found in YAC and BAC clones.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DB 1; Length 21;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Sequence 21 BP; 5 A; 5 C; 6 G; 5 T; 0 U; 0 Other;
```

The present invention describes a short interfering nucleic acid (siNA)

Example 3; SEQ ID NO 11; 139pp; English

New short interfering nucleic acid, useful e.g. for treatment and diagnosis of obesity or diabetes, downregulates expression of the stearoyl-CoA desaturase gene.

Thompson J;

Mcswiggen J, Beigelman L, (RIBO-) RIBOZYME PHARM INC.

WPI; 2003-721687/68.

2002US-0408378P. 2002US-0409293P. 2002US-0412304P. 2003US-0440129P.

2002US-0363124P. 2002US-0386782P. 2002US-0406784P

> 29-AUG-2002; 05-SEP-2002; 09-SEP-2002; 15-JAN-2003;

useful for identifying diagnostic and

13-FEB-2003; 2003WO-US004317.

20-FEB-2002;

WO2003070885-A2.

Synthetic.

28-AUG-2003

```
190 CTCCGGAGCCTCAGCC 205
                                                                                                                                                Conservative
                                                                                                                                             Similarity
                                                                                                                                               14;
                                                                                                                                           Query Match
                                                                                                                                              Local
                                                                                                                                                                RESULT 139
ADE27357/c
                                                                                                                                                Matches
g
                                                                                                                                                     ò
                                                                                                                                                  ö
                                                                                                                                                  Gaps
                                                                                                                                                  0
                                                                                                                                             0.4%; Score 16.2; DB 1
85.7%; Pred. No. 2e+02;
tive 0; Mismatches
```

```
..
0
that downregulates expression of the SCD (stearcy1-CoA desaturase) gene by RNA interference. Also described: (1) modulating expression of SCD genes in cells, tissue explants or organisms by introduction of SCD kits for in vitro or in vivo delivery of SiNA; (3) conjugates and/or complexes of SiNA; and (4) vectors that express siNA. SCD inhibiting siNAs have anorectic, antidiabetic, antiarteriosclerotic, cytostatic and virucide activities. The SiNAs can be used to modulate expression of SCD genes, in cells, tissue explants or organisms, e.g. for treating obseity; diabetes (types I and II); atherosclerosis; cancer and viral infections. They can also be used for drug screening; diagnosis; target identification and validation; genetic engineering; pharmacogenomics; studying gene function and gene mapping (e.g. of single-nucleotide polymorphisms). The present sequence represents an SCD SiNA, which is used in the exemplification of the present invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.4%; Score 16; DB 1; Length 19; 87.5%; Pred. No. 1.8e+02; tive 2; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 19 BP; 2 A; 10 C; 5 G; 0 T; 2 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADE27357 standard; RNA; 19 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADE27357;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             SXXX
```

```
The present invention describes a short interfering nucleic acid (siNA) that downregulates expression of the SCD (stearoyl-CoA desaturase) gene by RNA interference. Also described: (1) modulating expression of SCD genes in cells, tissue explants or organisms by introduction of siNA; (2) kits for in vitro or in vivo delivery of siNA; (3) conjugates and/or complexes of siNA; and (4) vectors that express siNA; SCD inhibiting siNAs have anorectic, antidiabetic, antiatteriosclerotic, cytostatic and virucide activities. The siNAs can be used to modulate expression of SCD genes, in cells, tissue explants or organisms, e.g. for treating obesity; diabetes (types I and II); atherosclerosis; cancer and viral infections. They can also be used for drug screening; diagnosis; target identification and validation; genetic engineering; pharmacogenomics; studying gene function and gene mapping (e.g. of single-nucleotide polymorphisms). The present sequence represents an SCD siNA, which is used in the exemplification of the present invention.
                                                                       short interfering nucleic acid; siNA; downregulation; inhibition; SCD; stearoyl-CoA dematurame; RNA interference; anorectic; antidiabetic; antiatreliosclerotic; cytostatic; virucide; obesity; diabetes; atherosclerosis; cancer; viral infection; drug screening; genetic engineering; pharmacogenomic; gene mapping; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     New short interfering nucleic acid, useful e.g. for treatment and diagnosis of obesity or diabetes, downregulates expression of the stearoyl-CoA desaturase gene.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ;
0
                                     Stearoyl-CoA desaturase siNA oligonucleotide SEQ ID NO:301.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     0.4%; Score 16; DB 1; Length 19;
100.0%; Pred. No. 1.8e+02;
tive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 19 BP; 2 A; 5 C; 10 G; 0 T; 2 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Example 3; SEQ ID NO 301; 139pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Mcswiggen J, Beigelman L, Thompson J;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    205
                                                                                                                                                                                                                                                                                                          13-FEB-2003; 2003WO-US004317.
                                                                                                                                                                                                                                                                                                                                                                11-MAR-2002; 2002US-0363124P.
06-JUN-2002; 2002US-0386782P.
29-AUG-2002; 2002US-0406784P.
                                                                                                                                                                                                                                                                                                                                                                                                                          05-SEP-2002; 2002US-0408378P.
09-SEP-2002; 2002US-0409293P.
20-SEP-2002; 2002US-041304P.
15-JAN-2003; 2003US-0440129P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AAT77163 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (RIBO-) RIBOZYME PHARM INC.
29-JAN-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               24-OCT-1997 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Local Similarity 100.
nes 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WPI; 2003-721687/68.
                                                                                                                                                                                                                              WO2003070885-A2
                                                                                                                                                                                                                                                                    28-AUG-2003
                                                                                                                                                                                             Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    190
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      16
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AAT77163;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 140
AAT77163/C
ID AAT771(XX
AC AAT771(XX
DT 24-OCT-
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Best Loca
Matches
 8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      셤
```

ö

```
PCR primers (AAT61331 and AAT77158-86) were designed for amplification of the human Batten disease CLN3 gene (see also AAT61306) exons 1-15. The CRC primers for exon 4 are given in AAT77163 and AAT77164. Novel mutations (see also AAT61332-88) have been discovered in the CLN3 gene of Bd patients using a combination of PCR, single strand conformation
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Batten disease polypeptide - useful to correct absence of wild type polypeptide, or as agonist to enhance activity of wild type polypeptide.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          se; antisense; casein kinase2-beta; mouse; antisense gene therapy;
cytostatic; antidiabetic; antiinflammatory; diabetes; cancer; tumour;
hyperproliferative disorder; breast cancer; prostate cancer;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Mole SE;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   5-methylcytidines"
                                                                                                          gene therapy; polymerase chain reaction; PCR; primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Mouse casein kinase 2-beta antisense oligonucleotide #14.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Indels
                                                                               diagnosis;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Breuning MH, Gusella JF,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   /*tag= b
/mod_base= OTHER
/note= "Phophorothioate backbone"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 20 BP; 6 A; 3 C; 8 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Score 16; DB 1; Lo
Pred. No. 1.9e+02;
                                                                               CLN3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     polymorphism analysis and direct sequencing
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /mod_base= OTHER
/note= "All cytidines are
                                                                            ceroid lipofuscinosis;
                       Batten disease gene exon 4 PCR primer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Disclosure; Page 31; 94pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.4%; Scc.
100.0%; Pre
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                2274 CAGCTCTGCCTTCACT 2289
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ABS65155 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                    96WO-US013896
                                                                                                                                                                                                                                                                                                                                                                                          95US-0003030P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        19 cagcrcrcccrrcacr 4
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Taschner PEM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ď
                                                                                                                                                                                                                                                                                                                                                                                                                                               (GEHO ) GEN HOSPITAL CORP (UYLE-) RIJKSUNIV LEIDEN.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ...20
*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 1997-179265/16
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Local Similarity
                                                                               Batten disease;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              modified base
                                                                                                                                                                                                                                                                                                                                    30-AUG-1996;
                                                                                                                                                                                                                                                                                                                                                                                             31-AUG-1995;
                                                                                                                                                                                                                    WO9708308-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Lerner TJ,
Gardiner MR;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         liver cancer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Mus musculus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          16;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   15-NOV-2002
                                                                                                                                                                                                                                                                            06-MAR-1997
                                                                                                                                                               Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ABS65155
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RESULT 141
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ABS65155
SX CC CC CC CX S X L L X X B B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B X X B
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      I I I I I I I I I I I X S X M M M X E X E X Y X Y I
```

```
RESULT
ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              The invention relates to a compound that is 8 - 50 nucleobases in length targeted to a nucleic acid molecule encoding Casein kinase 2-beta, and which specifically hybridises with and inhibits the expression of Casein kinase 2-beta, or which specifically hybridises with an 8-nucleobase portion of an active site on a nucleic acid molecule encoding Casein kinase 2-beta. Also included are: (1) a composition comprising the compound, and a carrier or diluent; (2) inhibiting the expression of casein kinase 2-beta in cells or tissues by contacting the expression of is inhibited, and (3) treating an animal having a disease or condition associated with Casein kinase 2-beta by administering to the animal the new compound so that the expression of Casein kinase 2-beta and for treating diseases or conditions associated with casein kinase 2-beta by administering the expression of Casein kinase 2-beta is inhibited. The antisense compounds are useful for modulating the expression of Casein kinase 2-beta is inhibited. The artisense compounds are useful for modulating the expression of Casein kinase 2-beta is inhibited. Whyerproliferative disorders, particularly cancer, such as breast cancer, prostate cancer, or liver cancer. The antisense compounds are also useful for diagnostics, therapeutics, prophylaxis, e.g. to prevent or delay infection, inflammation or tumour formation, as research reagents and kinds of various manner of various members of kits, and in distinguishing petween functions of various members of is a particus members of a particus members of propertices.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                               New antisense oligonucleotides targeted to nucleic acid encoding Casein kinase 2-beta, useful in diagnostic and research applications, or for treating a disease or condition associated with the expression of Casein kinase 2-beta.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ical pathway. The present sequence is an antisense oligonucleotide invention targeting mouse casein kinase 2-beta
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0.4%; Score 16; DB 1; Length 20; 100.0%; Pred. No. 1.98+02; Live 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human biallelic polymorphic marker upstream primer #210.
                                                                /*tag= d
/mod_base= OTHER
/note= "2'-methoxyethyl residues"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 6 A; 6 C; 6 G; 2 T; 0 U; 0 Other;
          /*tag= c
/mod_base= OTHER
/note= "2'-methoxyethyl residues'
                                                                                                                                                                                                                                                                                                                                                                                              Claim 3; Page 94; 142pp; English.
                                                                                                                                                                                                                                                                    Wyatt JR;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       CTGCTCTAGAGGGCCT 1003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      踞.
                                                                                                                                                                                 31-JAN-2002; 2002WO-US003159.
                                                                                                                                                                                                             08-FEB-2001; 2001US-00780175
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AAX09330 standard; DNA; 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           24-MAR-1999 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                CTGCTCTAGAGGGCCT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Local Similarity 100.
ses 16; Conservative
                                                    16. .20
                                                                                                                                                                                                                                        (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                    Mckay R, Freier SM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            biological pathway.
                                                                                                                                                                                                                                                                                             WPI; 2002-643409/69
                                                                                                                           WO200262954-A2
modified base
                                                      modified base
                                                                                                                                                      15-AUG-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       988
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 16
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AAX09330;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT 142
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Matches
ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      BXXXXX
```

```
solation of various biallele-specific oligonuclectide primers used in the isolation of various biallelic polymorphic markers found in the human genome (represented in AAX10269-X12937). These primers can be used in a method for determining polymorphic forms in an individual for used in e.g. of sagammaglobulinemia, diabetes insiplidus, Lesch-Nyhan syndrome, nuscular dystrophy, Wiskott-Aldrich syndrome, Fabry's disease, familial opposterolemia, polycystic kidney disease, hareditary herencoytosis, von Willebrand's disease, tubercous sclerosis, hereditary haemorrhagic telangiectesia, familial colonic polyposis, Ehlers-Danios syndrome, osteogenesis imperfecta, acute intermittent porphyria, syndrome, inflammation, cancer, diseases of the nervous system, infection by pathogenic microorganisms, and characteristics such as longevity, appearance (e.g. baldness, obesity), strength, speed, condust or therapeutic treatments. The isolated polymorphic nucleic acid segments can also be used to produce medicaments for the treatment or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Human, PCR, primer, vulnerary, anti-tumour, antirheumatic, antiarthritic, antiarteriosclerotic, cytostatic, neointima, scar, plaque, blood vessel; Toll-like receptor 4; adventitial cell; Tlr-4; ss.
Polymorphism; biallelic, human; forensic; paternity testing; disease; detection; phenotypic typing; characteristic; infection; hereditary; autoimmune disease; cancer; inflammation; drug; therapy; medicament; treatment; marker; primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             New isolated nucleic acid segments from the human genome - used for determining polymorphic forms for use in e.g. forensics, paternity testing or phenotypic typing for disease.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15.8; DB 1; Length 19; 89.5%; Pred. No. 1.9e+02; ive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human Toll-like receptor 4, Tlr-4, PCR primer #1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 19 BP; 5 A; 7 C; 5 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (WHED ) WHITEHEAD INST BIOMEDICAL RES.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Claim 15; Page 71; 310pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  186 CAGACTCCGGAGCCTCAGC 204
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1 CAGACTCTGGAGCCACAGC 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Hudson T;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ACC70795 standard; DNA; 19 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                  97WO-US020313
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          96US-0030455P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    prophylaxis of such diseases
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Local Similarity 89.5
hes 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Lander ES, Wang D,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WPI; 1998-286974/25.
                                                                                                                                                                                                                                Homo sapiens
                                                                                                                                                                                                                                                                                                 WO9820165-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                  05-NOV-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          06-NOV-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       20-NOV-2003
                                                                                                                                                                                                                                                                                                                                                                          14-MAY-1998.
                                                                                                                                                                                      Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACC70795;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Matches
```

Camara

Homo sapiens. EP1302206-A1

16-APR-2003

```
The invention relates to a novel process for increasing the metabolites of carotene content of a plant. The novel process comprises upregulating at least one gene which encodes carotene hydroxylase activity. The compositions of the novel process have lycopene beta-cyclase or a beta-carotene hydroxylase activity. The process is useful for increasing the metabolites of carotene content of a plant, comprising transforming a plant cell from which viable plants may be recovered, using a plant expression cassette, or a DNA construct, and generating viable plants from the cell. The carotene metabolites are useful for increasing zeaxanthin and beta-carotene, including oxygenated carotenoids. This polymucleotide sequence represents an RT-PCR primer used in the process for the expression of the introduced proteins and endogenous carotenoid genes of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        human; 88; SiRNA; short interfering nucleic acid; siNA;
protein tyrosine phosphatase-1B; PTP-1B; RNA interference; RNAi;
micro-RNA; miRNA; short hairpin RNA; shRNA; gene silencing; antisense;
obesity; insulin resistance; diabetes; anorectic; antidiabetic;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Antisense siNA that down regulates human PTP-1B expression (SeqID 279)
                                                                                    Increasing the metabolites of carotene content in a plant useful for producing recombinant plants comprises upregulating a gene encoding carotene hydroxylase activity.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      0.4%; Score 15.8; DB 1; Length 19; 89.5%; Pred. No. 1.9e+02; ive 0; Mismatches 2; Indels
Dharmapuri S, Pallara P,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 19 BP; 2 A; 6 C; 3 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                             Example 1; Page 12; 21pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 1716 TCACTCTCCAGTCTTCAGG 1734
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            rcrcrrrccagrcrrcage 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      2002US-0386782P.
2002US-0020670S.
2002US-0406784P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADF75738 standard; RNA; 19 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    11-FEB-2003; 2003WO-US004123
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              2002US-0358580P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          2002US-0408378P.
2002US-0409293P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         2003US-0440129P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Mcswiggen J, Beigelman L,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (RIBO-) RIBOZYME PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 26-FEB-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Best Local Similarity 89.5
Matches 17, Conservative
Giuliano G, Rosati C,
                                          WPI; 2003-714401/68.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WPI; 2003-697604/66
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WO2003070881-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        06-JUN-2002; 26-JUL-2002; 29-AUG-2002; 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20-FEB-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               05-SEP-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    09-SEP-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         15-JAN-2003;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        11-MAR-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         28-AUG-2003.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      antisense.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ADF75738;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 145
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ADF75738/c
ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       metabolite, carotene, plant, carotene hydroxylase, lycopene beta-cyclase, beta-carotene hydroxylase, zeaxanthin, beta-carotene; oxygenated carotenoid, RT-PCR, primer, carotenoid, 88.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  The present invention relates to a method for interfering with the formation of a neointima/scar and/or a plaque in a blood vessel by providing a ligand capable of modulating Toll-like receptor activity of adventitial cells. The method is useful for reducing the formation of a neointima/scar and/or a plaque in a blood vessel after stenting, angioplasty, heart transplantation, by pass surgery, arteriovenous shunting and infection, especially bacterial infection. The method is also useful for modulating tumour growth, and for modulating the effects of rheumatoid architis. The present sequence is a PCR primer for human
                                                                                                                                                                                                                                                                                                                                                                              Interfering with the formation of a neointima/scar and/or a plaque in a blood vessel, useful for modulating tumor growth, comprises providing a ligand capable of modulating Toll-like receptor activity of adventitial cells.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.4%; Score 15.8; DB 1; Length 19; 89.5%; Pred. No. 1.9e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Endogenous carotenoid gene expression RT-PCR primer #12
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 19 BP; 3 A; 8 C; 2 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (CNEN ) ENEA ENTE NUOVE TECNOLOGIE ENERGIA.
(BIOJ ) BIOGEN SRL.
                                                                                                                                                                                                                   (UYUT-) UNIV UTRECHT MEDISCH CENT. (UYUT-) RIJKSUNIV UTRECHT.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Disclosure; Page 7; 23pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               2273 TCAGCTCTGCCTTCACTAC 2291
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1 rcascrcrscrrcracrac 19
                                                                                                                              11-OCT-2001; 2001EP-00203846
                                                                                                                                                                         11-OCT-2001; 2001EP-00203846.
                                                                                                                                                                                                                                                                                       De Kleijn DPV, Pasterkamp G;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       08-NOV-2002; 2002EP-00425681.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   09-NOV-2001; 2001IT-RM000670
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Toll-like receptor 4 (Tlr-4)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADE78588 standard; DNA; 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    17; Conservative
                                                                                                                                                                                                                                                                                                                                    WPI; 2003-484923/46.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Best Local Similarity
```

Query Match

Matches

29-JAN-2004

ADE78588;

RESULT 144

g

Unidentified EP1323825-A2

02-JUL-2003

Usman N;

ö

Gaps

```
11-MAR-2002; 2002US-0563124P.
06-JUN-2002; 2002US-0386782P.
26-JUL-2002; 2002US-0406784P.
29-AUG-2002; 2002US-0406784P.
05-SEP-2002; 2002US-0408378P.
09-SEP-2002; 2002US-0409293P.
15-JAN-2003; 2003US-041029P.
                                                                                                                                                                                                                                                        11-FEB-2003; 2003WO-US004123
                                                                                                                                                                                                                                                                 20-FEB-2002; 2002US-0358580P
                                                                                                                                                                                                                                       WO2003070881-A2
                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                28-AUG-2003.
                                                                                                                                                                              ADF75553;
                                                                                                                      Query Match
                                                                                                                              Matches
                                                                                                                                                             RESULT 146
                                                                                                                                                                 ð
                                                                                                                                               셤
```

Mcswiggen J, Beigelman L, Usman N;

WPI; 2003-697604/66

(RIBO-) RIBOZYME PHARM INC.

This invention relates to novel short interfering nucleic acid (siNA)

molecules that downregulate expression of a protein tyrosine phosphataseIB (PTP-1B) gene by RNA interference (RNAi). Specifically, the siNAs can
be short interfering RNA interference (RNAi). Specifically, the siNAs can
cof PTP-1B. The present invention describes sequence-specific posttrancriptional gene silencing in animals using siNA molecules and
antisense oligonucleotides to modulate PTP-1B gene expression or
crivity. Furthermore, these siNA molecules provide useful reagents for a
crivity of therapeutic and diagnostic purposes, and as such can be used
for treating obseity, insulin resistance or diabetes (types I and II), as
well as for drug screening, target identification and validation, genetic
engineering, pharmacogenomics and for studying gene function and gene
mapping (for example of single-nucleotide polymorphisms). Accordingly,
these molecules exhibit anorectic and antidiabetic activities. ö sequence is an antisense siNA molecule that targets human human; 88; siRNA; short interfering nucleic acid; siNA; protein tyrosine phosphatase-1B; PTP-1B; RNA interference; RNAi; micro-RNA; miRNA; short hairpin RNA; shRNA; gene silencing; antisense; obesity; insulin resistance; diabetes; anorectic; antidiabetic. Gaps New short interfering nucleic acid, useful e.g. for treatment and diagnosis of obesity, downregulates expression of a protein tyrosine phosphatase-18 gene. Sense siNA that down regulates human PTP-1B expression (SeqID 94) ; 0 Match 0.4%; Score 15.8; DB 1; Length 19; Local Similarity 89.5%; Pred. No. 1.9e+02; les 17; Conservative 0; Mismatches 2; Indels Sequence 19 BP; 7 A; 3 C; 7 G; 0 T; 2 U; 0 Other; Example 3; SEQ ID NO 279; 140pp; English. 3162 TGCCCCTTCCATTTTAAGT 3180 19 récécérrécactirgagi 1 ВР oligonucleotide sequence is PTP-1B RNA of the invention. ADF75553 standard; RNA; 19 26-FEB-2004 (first entry)

```
This invention relates to novel short interfering nucleic acid (siNA)
molecules that downregulate expression of a protein tyrosine phosphatase-
18 (PTP-1B) gene by RNA interference (RNAi). Specifically, the siNAs can
be short interfering RNA (siRNA), double-stranded RNA (dsRNA), micro-RNA
(miRNA) or short hairpin RNA (shRNA), all of which can mediate inhibition
c f PTP-1B. The present invention describes sequence-specific post-
trancriptional gene silencing in animals using siNA molecules and
antisense oligonucleotides to modulate PTP-1B gene expression or
activity. Furthermore, these siNA molecules provide useful reagents for a
variety of therapeutic and diagnostic purposes, and as such can be used
for treating obesity, insulin resistance or diabetes (types I and II), as
well as for drug screening, target identification and validation, genetic
mapping (for example of single-nucleotide polymorphisms). Accordingly,
these molecules exhibit anorectic and antidiabetic activities. This
oligonucleotide sequence is a sense siNA molecule that targets human PTP-
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RNA interference; short interfering nucleic acid; siNA; short interfering RNA; siRNA; double-stranded RNA; micro-RNA; miRNA; short hairpin RNA; shrNA; expression modulation; gene therapy; drug screening; diagnosis; therapeutic target identification; pharmacogenomics; gene function analysis; gene mapping; tumour necrosis factor; TNR; human; DNA-RNA hybrid; ss, antibacterial; immunosuppressive; antirheumatic; antiarthritic; anti-HIV; antipsoriatic; antinflammatory; septic shock; rheumatoid arthritis; HIV/AIDS; psoriasis; inflammation; autoimmune disease; target sequence.
New short interfering nucleic acid, useful e.g. for treatment and diagnosis of obesity, downregulates expression of a protein tyrosine phosphatase-1B gene.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.4%; Score 15.8; DB 1; Length 19; 22.6%; Pred. No. 1.9e+02; ve 7; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 19 BP; 2 A; 7 C; 3 G; 0 T; 7 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human TNF sinA oligonucleotide SEQ ID NO:66.
                                                                                                 Example 3; SEQ ID NO 94; 140pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               3162 TGCCCCTTCCATTTTAAGT 3180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1 UGCCCCUUCCACUUUGAGU 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ADG34714 standard; RNA; 19 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         2002US-0363124P.
2002US-0386782P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            2002US-0406784P.
2002US-0408378P.
2002US-0409293P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  09-SEP-2002; 2002US-0409293P.
28-NOV-2002; 2002US-0429359P.
15-JAN-2003; 2003US-0440129P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20-FEB-2003; 2003WO-US004741.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2002US-0358580P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        52.6%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   26-FEB-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Best Local Similarity 52.6
Matches 10, Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                        1B RNA of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WO2003070897-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         06-JUN-2002;
29-AUG-2002;
05-SEP-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       20-FEB-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         11-MAR-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           28-AUG-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ADG34714;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 147
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ADG34714/
      ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     g
```

10001863-3.81.rng

WO2003070897-A2

```
The invention relates to short interfering nucleic acids (siNA) which downregulate expression of the human tumour necrosis factor (TNF) gene by RNA interference. The siNAs may or may not comprise ribonucleotides and may be double or single stranded. They further comprise sense and autisense regions, or alternatively are assembled from a sense coligonucleotide short interfering RNA (siRNA), double-stranded RNA, micro-RNA include short interfering RNA (siRNA), double-stranded RNA, micro-RNA (miRNA) and short hairpin RNA (siRNA). The siNAs can be unmodified or chemically synthesised, expressed from a vector or enzymatically chemically synthesised. The invention also relates to kits for the in vitro or in control of siNA, conjugates and/or complexes of siNA, and vectors that express siNA. The siNAs are used to modulate expression of the TNF gene in cells, tissue explants or organisms (e.g., by ex vivo gene therapy), or in grafts and transplants for the treatment of a variety of conditions. The TNF siNAs have antibacterial, immunosuppressive, antiarthritis, antiarthritic, anti-HIV, antipsoriatic and autoimmune diseases. The siNAs are also useful for drug screening, diagnosis, theumatoid arthritis, HIV/AIDS, psoriasis, inflammation and autoimmune diseases. The siNAs are also useful for drug screening, dance also useful for drug screening, dance also useful for drug screening, studying gene function, and gene mapping (e.g., of single nucleotide polymorphisme). The present sequence represents the mine and autoimmune control of a human TNF-targeted double-stranded sinA, which is
                                                                                                                                                                              New short interfering nucleic acid, useful e.g. for treatment and diagnosis of septic shock or rheumatoid arthritis, downregulates expression of the tumor necrosis factor gene.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          identical to the TNF transcript target sequence
                                                                                                                                                                                                                                                                                    Example 3; SEQ ID NO 66; 141pp; English
                        (RIBO-) RIBOZYME PHARM INC
                                                                             Beigelman L;
                                                                                                                               WPI; 2003-697609/66.
                                                                             Mcswiggen J,
```

Gaps ö 0.4%; Score 15.8; DB 1; Length 19; 89.5%; Pred. No. 1.9e+02; rative 0; Mismatches 2; Indels Sequence 19 BP; 4 A; 5 C; 3 G; 0 T; 7 U; 0 Other; Best Local Similarity 89.5 Matches 17; Conservative Query Match Matches

ö

3414 TTTCAAGGAAGTATGGAAA 3432 rcrcaaggaagrcrggaaa 1 13 ઠ a

ADG34802 standard; RNA; 19 BP ADG34802; RESULT 148 ADG34802 

26-FEB-2004 (first entry)

Human TNF sinA oligonucleotide SEQ ID NO:154.

pharmacogenomics; gene function analysis; gene mapping; tumour necrosis factor; TNF; human; DNA-BNA hybrid; ss; antibacterial; fimunosoppressive; antibarheumatic; antiarthritic; anti-HIV; antibaciatic; antiinflammatory; septic shock; rheumatoid arthritis; HIV/AIDS; psoriasis; inflammation; autoimmune disease. RNA interference; short interfering nucleic acid; siNA; short interfering RNA; siRNA; double-stranded RNA; micro-RNA; miRNA; short hairpin RNA; shRNA; expression modulation; gene therapy; drug screening; diagnosis; therapeutic target identification;

Homo sapiens. Synthetic

Human apolipoprotein B (ApoB) oligonucleotide segid 3026.

16-DEC-2004 (first entry)

ADR78541;

3541/c ADR78541 standard; DNA; 19 BP.

ADR78541/

```
The invention relates to short interfering nucrous since in the invention relates to short interference. The sinks may or may not comprise ribonucleotides and may be double or single stranded. They further comprise sense and antisense regions, or alternatively are assembled from a sense and configuration of the sinks of sense and antisense oligonucleotide. Specifically, the sinks include short interfering RNA (siRNA). The sinks can be unmodified or chemically modified, can contain deoxyribonucleotides, and can be chemically synthesised, expressed from a vector or enzymatically contain an also relates to kits for the in vitro or in vivo delivery of sink, conjugates and/or complexes of sink, and vectors that express sink. The sinks are used to modulate expression of the TNF gene in cells, tissue explants or organisms (e.g., by ex vivo gene to conditions. The TNF sinks are used to modulate expressive, antirheumatic, antiarthritic, anti-HIV, antipsoriatic and anticheumatic antiarthritis, They may be used for treating septic shock, theumatoid arthritis, HUVAIDS, psoriasis, inflammation and autoimmune diseases. The sinks are also useful for drug screening, diagnosis, therapeutic target identification and validation, genetic entire treating septic of the therapeutic target identification and validation, genetic entire.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            pharmacogenomics, studying gene function, and gene mapping (e.g., of single nucleotide polymorphisms). The present sequence represents the lower strand of a human TNF-targeted double-stranded siNA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                         New short interfering nucleic acid, useful e.g. for treatment and diagnosis of septic shock or rheumatoid arthritis, downregulates expression of the tumor necrosis factor gene.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              The invention relates to short interfering nucleic acids (siNA)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Score 15.8; DB 1; Length 19;
Pred. No. 1.9e+02;
4; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 19 BP; 7 A; 3 C; 5 G; 0 T; 4 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Example 3; SEQ ID NO 154; 141pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      3414 TTTCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1 ucuchaggaagucuggaah 19
                                                                                                                                                                06-JUN-2002; 2002US-0386782P
29-AUG-2002; 2002US-0406784P
05-SEP-2002; 2002US-0406378P
09-SEP-2002; 2002US-0409293P
28-NOV-2002; 2002US-04293S9P.
                                                                                  20-FEB-2003; 2003WO-US004741.
                                                                                                                                                                                                                                                                                15-JAN-2003; 2003US-0440129P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match 0.4%;
Best Local Similarity 68.4%;
                                                                                                                                                                                                                                                                                                                           (RIBO-) RIBOZYME PHARM INC
                                                                                                                                                                                                                                                                                                                                                                       Mcswiggen J, Beigelman L;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            13; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2003-697609/66.
                                         28-AUG-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 149
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Matches
g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ઠે
```

Sequence 19 BP; 9 A; 5 C; 1 G; 4 T; 0 U; 0 Other; can be used to control ApoB gene expression.

SXS

```
WPI; 2004-677362/66.
                                 WO2004080406-A2.
                                                                  25-APR-2003;
25-APR-2003;
09-MAY-2003;
                                                                                                    Manoharan M,
                            Homo sapiens.
                                                           13-MAR-2003;
                                                                           08-AUG-2003;
                                                                              11-AUG-2003;
                                                  07-MAR-2003;
                                                                17-APR-2003;
                                                                                         07-NOV-2003;
                                                                                    09-OCT-2003;
                                                     12-MAR-2003;
                                                        13-MAR-2003;
                                                             14-APR-2003;
                                                                                 26-SEP-2003;
                                       23-SEP-2004
```

artery cytostatic; anticonvulant; notropic; muscula, anti-HIV; what interference; iRNA; antisense technology; lipid metabolism; cholesterol imbalance; dyslipidaemia hypercholesterolaemia; coronary attery disease; CAD; coronary heart disease; CHD; atherosclerosis; hepatic glucose production; glucose-metabolism-related disorder; disbates; cancer; breast cancer; colon cancer; lung cancer; neurological disease; Huntington disease; spinocerebellar ataxia; viral disease; AIDS; apolipoprotein B; apoB; ss. antilipemic; cardiant; vasotropic; antiarteriosclerotic; antidiabetic; Interference RNA agent useful for treating dyslipidemias, coronary disease, diabetes, cancer or neurological disease, comprises sense sequence and antisense sequence which has specific modifications. Example 5; SEQ ID NO 3026; 378pp; English. 2003US-0465665P 2003US-0465802P 2003US-0493986P 2003US-0494597P 2003US-0506341P 2003US-0510246P 2003US-0455050P. 2003US-0462894P. 2003US-0463772P. 2003US-0454265P. 2003US-0454962P. 2003US-0510318P. 2003US-0518453P. 08-MAR-2004; 2004WO-US007070 Bumcrot D; (ALNY-) ALNYLAM PHARM.

The invention describes a RNA interference (iRNA) agent (I) comprising a sense sequence and an antisense sequence, where the sense sequences have one or more asymmetrical 2-O alkyl modifications, the antisense sequence and the antisense sequence asymmetrical phosphorothicate modifications and the antisense sequence targets a human gene sequence. Also described and the antisense sequence targets a human gene sequence. Also described care: a pharmaceutical preparation comprising (I); reducing (II); involves selecting a sequence with activity and stabilising (I), involves selecting a sequence with activity and consortivity; a kit comprising (I) and instruction for its use; and a device the modification decreases nuclease sensitivity while not decreasing its activity; a kit comprising (I) and instruction for its use; and a device the modification decreases nuclease sensitivity while not decreasing its activity; a kit comprising (I) and instruction for its use; and a device that can be dispense or administer a composition comprising (I). (I) is useful for reducing apoB-100 levels or glucose-6-phosphatase levels.

Cueseful for reducing apoB-100 levels or glucose-6-phosphatase levels.

Cuevals of cholesterol, and/or disregulation of lipid metabolism. The disorder is choses for coronary artery disease (CHD), coronary heart cype-companial publication and atherosclerosis. (I) is administered to a subject to inhibit hepatic glucose production or for treating glucose-metabolism.

Cueseful disorder e.g. diabetes or type-2 diabetes. (I) is useful for relating the diseases as mentioned above, cancer (e.g. breast) or coronary artery disease (c.g., Hung cancer), neurological

ö Interference RNA agent useful for treating dyslipidemias, coronary artery disease, diabetes, cancer or neurological disease, comprises sense sequence and antisense sequence which has specific modifications. The invention describes a RNA interference (iRNA) agent (I) comprising a sense sequence and an antisense sequence, where the sense sequences have one or more asymmetrical 2'-O alkyl modifications, the antisense sequences have one or more asymmetrical phosphorothicate modifications antilipemic; cardiant; vasotropic; antiarteriosclerotic; antidiabetic; cytostatic; anticonvulsant; nootropic; muscula; anti-HIV; MAX interference; iRNA; antisense technology; lipid metabolism; cholesterol imbalance; dyslipidaemia hypercholesterolaemia; coronary artery disease; CAD; coronary heart disease; CHD; atherosclerosis; hepatic glucose production; glucose-metabolism-related disorder; diabetes; cancer; breast cancer; colon cancer; lung cancer; neurological disease; Huntingron disease; spinocerebellar ataxia; viral disease; AIDS; apolipoprotein B; apoB; ss. Gaps .. 0 0.4%; Score 15.8; DB 1; Length 19; 89.5%; Pred. No. 1.9e+02; ive 0; Mismatches 2; Indels Human apolipoprotein B (ApoB) oligonucleotide segid 408. Example 5; SEQ ID NO 408; 378pp; English. 3699 AGATGTTTTTTTTCAG 3717 19 AGATGGTTAGTTTTTTCAG 1 2003US-0454265P. 2003US-0454962P. 2003US-045496P. 2003US-0462894P. 2003US-0463472P. 2003US-0463665P. 2003US-0506341P. 2003US-0510246P. 2003US-0510318P. 2003US-0518453P. ВР 2003US-0469612P. 2003US-0493986P. 2003US-0494597P. 08-MAR-2004; 2004WO-US007070 2003US-0452682P ADR75923 standard; DNA; 19 (first entry) Conservative Manoharan M, Bumcrot D; (ALNY-) ALNYLAM PHARM. WPI; 2004-677362/66. Similarity WO2004080406-A2. 17-APR-2003; 2 25-APR-2003; 2 Homo sapiens. 14-APR-2003; 25-APR-2003; 09-MAY-2003; 07-MAR-2003; 13-MAR-2003; 13-MAR-2003; 08-AUG-2003; 11-AUG-2003; 26-SEP-2003; 16-DEC-2004 23-SEP-2004. 17; ADR75923; Query Match Local RESULT 150 ADR75923/c Matches ð

```
are: a pharmaceutical preparation comprising (II); reducing (MI) apoB-100 levels or glucose-6-phosphatase levels in a subject; producing (II); stabilising (II); involves selecting a sequence with activity and introducing one or more asymmetrical modification in the sequence, where the modification decreases nuclease sensitivity while not decreasing its activity; a kit comprising (I) and instruction for its use; and a device that can be dispense or administer a composition comprising (I). (I) is useful for reducing apoB-100 levels or glucose-6-phosphatase levels. The subject is suffering from a disorder characterised by elevated or otherwise unwanted expression of apoB-100, elevated or otherwise unwanted expression of apoB-100, elevated or otherwise unwanted characterised by elevated or otherwise unwanted expression of apoB-100, clevated or otherwise unwanted characterised by elevated or otherwise unwanted expression of apoB-100, clevated or otherwise unwanted characterised by closesterolaemia, and/or disregulation of lipid metabolism. The dispense from the HDL/LDL cholesterol imbalance, dyslipidaemias, hypercholesterorolaemia, statin-resistant clisease (CHD) and atherosclerosis. (I) is administered to a subject to inhibit hepatic glucose production or for treating glucose-metabolism. Creating the diseases as mentioned above, cancer (e.g. breast, colon or lung cancer), neurological disease (e.g., Huntington disease or lung cancer), neurological disease (e.g., Huntington disease or lung cancer elected librated or elected elect
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        spinocerebellar ataxia) or viral disease (e.g., AIDS). This sequence represents a human apolipoprotein B (ApoB) antisense oligonucleotide that
  and the antisense sequence targets a human gene sequence. Also described
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /*tag= a
/note= "each base is linked by N3'-P5' phosphoramidate
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Oligodeoxyribonucleotide; intersubunit linkage; phosphoramidate intersubunit; antisense activity; nuclease resistant; in-vitro cell growth inhibition assay; infection; smooth muscle cell proliferation disorder; inflammatory process; genetic disorder; cancer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Modified oligonucleotide containing N3'-P5' phosphoramidates
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Score 15.8; DB 1; Length 19;
Pred. No. 1.9e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 19 BP; 9 A; 5 C; 1 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            can be used to control ApoB gene expression.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Chen J;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      3699 AGATGTTTTTTTTCAG 3717
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 19 AGATGGTTAGTTTTTTTTAG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (LYNX-) LYNX THERAPEUTICS INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAX59720 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             94US-00214599.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       95WO-US003575
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      94US-00210505
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Schultz RG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        22-JUL-1999 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Local Similarity 89.5
nes 17, Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          linkages"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         œ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gryaznov SM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WO9525814-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20-MAR-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    18-MAR-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             18-MAR-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        28-SEP-1995.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AAX59720;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Best Loca
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 151
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AAX59720
g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ò
```

```
disorders, inflammatory processes, certain genetic disorders, cancers, etc. . The present sequence represents an oligonucleotide of the invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                               The specification describes oligodeoxyribonucleotides having contiguous nucleoside subunits joined by intersubunit linkages, where at least 3 contiguous subunits are joined by phosphoramidate intersubunits. The coligodeoxyribonucleotides has a sequence of mucleoside subunits effective to form a duplex with a target nucleic acid molecule. The oligodeoxyribonucleotides are more resistant to nuclease digestion and have improved RNA and dsDNA hybridisation characteristics, relative to oligonucleotides not containing N3'-P5' phosphoramidate linkages. They also have excellent antisense activity against complementary mRNA targets in .vitro cell growth inhibition assays. They also exhibit low cytcoxicity. They may be used in diagnostic and therapeutic applications, e.g., in combatting infections agents such as bacteria, viruses, etc. or in treatment of smooth muscle cell proliferation
                                Oligo:nucleotide N3'-P5' phosphoramidate(s) - have improved resistance toward phosphodiesterase digestion, and form stable duplexes with DNA and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Isolated nucleic acid encoding hepatocyte nuclear factor 1-alpha and beta - useful for detecting susceptibility for non-insulin dependent diabetes, especially maturity-onset diabetes of the young.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Hepatocyte nuclear factor 1 beta; HNF-1 beta; MODY4; human; transcription factor; maturity onset diabetes of the young; TCF2; diabetes; NIDDM; diagnosis; therapy; DCOH; PCBD; PCR; primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ŝ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Furuta H, Menzel
                                                                                                                                                                                                                                                                                                                                                                                                                                                       DB 1; Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 8 A; 0 C; 0 G; 12 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.4%; Score 15.8; DB 1
89.5%; Pred. No. 2e+02;
iive 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Kaisaki PJ,
                                                                                                    Disclosure; Page 55; 101pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      DCoH (PCBD) gene exon 4 PCR primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          3303 TATTTTTTTTTTATAT 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1 rarararrrrrrarar 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAV52722/c
ID AAV52722 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          oda N,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     97WO-US016037.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      96US-0028056P.
96US-0029679P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     96US-0025719P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Best Local Similarity 89.5
Matches 17, Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Yamagata K,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (ARCH-) ARCH DEV CORP.
WPI; 1995-344627/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 1998-271667/24.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WO9811254-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     10-SEP-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     21-DEC-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       .0-SEP-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      30-OCT-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 19-MAR-1998.
                                                                    RNA strands.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       12-OCT-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Horikawa Y;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAV52722;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Bell GI,
                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 152
 ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          셤
```

```
WO9949065-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                            SISI (-SISI)
                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                       26-MAR-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Miraglia LJ,
                                                                                                                                                                                                                                                             07-JAN-2000
                                                                                                                                                                                                                                                                                                                                       restenosis;
                                                                                                                                                                                                                                                                                                                                                                                                      30-SEP-1999
                                                                                                                                                                                                                                           AAZ37657;
                                                                                                                                                                                                                                                                                                                                                         Synthetic
8888888888888888
                                                                                                                                                                                                                                  ò
                                                                                                                                                                           엄
                                                                                                                               ö
                                                                                                                                                                                                                                                                                        human; B7; T cell; inflammation; autoimmune disease; cell activation;
                This is a reverse PCR primer designed for use with a forward primer (see AAV52212) in the PCR amplification of exon 4 of the human DCOM (PCBD) gene. The bifunctional DCOM/PCBD protein stabilises dimers of hepatocyte nuclear factor-1 beta (HNF-1 beta, see AAW71581) with itself or with HNF-1 alpha (see AAW71559). Mutations of HNF are associated with MODY (maturity onset diabetes of the young) type diabetes. No diabetes-sesociated mutations were found in DCOH
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              New oligo:nucleotide(s) that modulate expression of B7 proteins - used for, e.g. controlling activation and proliferation of T cells, particularly for treatment, diagnosis and prevention of inflammation.
                                                                                                                                                                                                                                                                                                                                                         1. .5
/*tag= a
/*total phosphorothioate linkages, optionally
modified with 2'fluoro-1 or 2'-methoxyethoxy"
                                                                                                                                                                                                                                                                                                                                                                                                                                                            /*tag= d
/note= "Optional phosphorothioate linkages, optionally
modified with 2'fluoro-1"
                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                          "optionally modified with 2'-methoxyethoxy"
                                                                                                                                .
0
                                                                                                            DB 1; Length 20;
                                                                                                                              2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                      *tag= b
'note= "Phosphorothioate linkages"
                                                                                           Sequence 20 BP; 4 A; 7 C; 4 G; 5 T; 0 U; 0 Other;
                                                                                                          Score 15.8; DB 1
Pred. No. 2e+02;
0; Mismatches
                                                                                                                                                                                                                                                                      Human B7-1 targetted oligonucleotide 12370.
Example 8; Page 146; 363pp; English.
                                                                                                                                                                                                                                                                                                                                               Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Example 1; Page 35; 120pp; English.
                                                                                                                                                3068 TGACTGAACTGGGTGTTCA 3086
                                                                                                                                                                  ~
                                                                                                            0.4%;
                                                                                                                                                                  rcacrcaacrccaacrcca
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     97WO-US023270.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      96US-00777266
                                                                                                                                                                                                                AAV48008 standard; DNA; 20
                                                                                                                                                                                                                                                    19-OCT-1998 (first entry)
                                                                                                                                17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                   . 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Bennett CF, Vickers TA;
                                                                                                                                                                                                                                                                                                                                                                                                                        14. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                 tag= c
                                                                                                                                                                                                                                                                                                                                                                                                                                           'note=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 1998-387783/33.
                                                                                                                                                                                                                                                                                                  cell proliferation.
                                                                                                                     Local Similarity
                                                                                                                                                                                                                                                                                                                                                 Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                       modified_base
                                                                                                                                                                                                                                                                                                                                                                                             modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                   modified_base
                                                                                                                                                                                                                                                                                                                               Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    16-DEC-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WO9829124-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      31-DEC-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  09-JUL-1998.
                                                                                                                                                                                                                                                                                                                     Synthetic
                                                                                                                                                                  20
                                                                                                                                                                                                                                  AAV48008;
                                                                                                             Query Match
                                                                                                                                                                                            ò
                                                                                                                                                                g
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AAZ37473-Z37738 represent human mdm2 phosphorothicate oligonucleotides. AAZ3741, AAZ37472, AAZ37739, AAZ37740 and AAZ37741 are used in the exemplification of the present invention. The present invention describes novel nucleotide antisense compounds, targetted to the 5' untranslated, translation termination codon, or 3' untranslated region of a nucleic acid encoding human mdm2, that modulates expression of human mdm2. The oligonucleotides mediate their effect by antisense inhibition of hyperproliferative gene expression. The antisense compound is used to treat an animal having a disease or condition associated with mdm2, especially of the blood, brain, breast, lung or soft tissue, or psoriasis, fibrosis, atherosclerosis or restenosis
The oligonucleotides which specifically hybridise to B7 modulate its expression (and thus T cell activation and proliferation). This is particularly useful for treatment and prevention of inflammation and autoimmune diseases, e.g. asthma, (juvenile) diabetes, myasthenia gravis, Grave's disease, rheumatoid arthritis, allograft rejection, psoriasis, (systemic) lupus erythematosus, multiple sclerosis, contact dermaticis, rhintis, allergy, cancer and metastases. The oligonucleotides may also be used to manipulate T cell activation ex vivo; to determine or detect B7 protein expression; for diagnosis; as assay and purification reagents, and to study physiological roles of B7 proteins
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          New antisense compounds used to treat eg. hyperproliferative conditions
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human mdm2 gene; proliferation; tumour; phosphorothioate; p53; cancer; antisense; modulation; oligonucleotide; expression; hihlbition; hyperproliferation; blood cancer; brain cancer; brain cancer; brain cancer; lung cancer; soft tissue cancer; psoriasis; fibrosis; atherosclerosis;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.4%; Score 15.8; DB 1; Length 20; 89.5%; Pred. No. 2e+02; ive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Cowsert LM;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Human mdm2 phosphorothioate oligodeoxynucleotide #187.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 0 A; 4 C; 4 G; 12 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 20 BP; 7 A; 4 C; 8 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Graham MJ, Monia BP,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Example 9; Page 52; 157pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    3766 recreedarcecreer 3784
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            99WO-US006702.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20 TGGCTGGCATCCCTCTCCT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      98US-00048810.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AAZ37657 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Nero P,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
Best Local Similarity
Local 17; Conserva
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WPI; 1999-610754/52.
```

Antisense oligonucleotide, phosphorothioate, TNFalpha, cytokine, inhibit, tumour necrosis factor alpha, inflammatory bowel disease, diabetes; rheumatoid arthritis, infections amountaiple sclerosis; hepatitis, pancreatitis, atopic dermatitis, allograft rejection; autoimmune disease;

inflammatory disease; ss.

AAX3667;

ઠ g WO200020645-A1.

Synthetic.

Human TNFalpha antisense oligonucleotide ISIS# 104785.

(first entry)

16-AUG-2000

AAA41139 standard; DNA; 20 BP.

```
AAA41139
                            This sequence represents a PCR primer used in the method of the invention. The method is for detecting the presence of alleles for invention. The method is for detecting the presence of alleles for glaucoma comprising haplotype analysis of human chromosome 2 and 6 respectively, where the haplotypes are associated with loci (GCIB and GLC6p25 respectively. The primers are used to amplify gene sequences to generate information necessary to compile haplotype profiles can be used to localise, isolate and symptomatic glaucoma. They can also be used to localise, isolate and identify the CGLC1B and GLC6p25 loci so that detection of individuals with glaucoma is enhanced. The haplotype analyses also provide means for identification and following of mutant alleles in pedigrees or populations. Cidentification of presymptomatic individuals using the methods allows intervention in the disease process and obviates the impact of inheriting a metant allele causing disease, by medically disrupting the initiation
                                             ö
                                                                                                                                                                                                                                                                                                                    PCR primer; detection; glaucoma allele; haplotype analysis; human; GLC1B; chromosome 2; chromosome 6; GLC6p25; haplotype profile; presymptomatic glaucoma; symptomatic glaucoma; ss.
                                             Gaps
                                             ö
             Length 20;
                                             Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Haplotype analyses for indirect detection of glaucoma
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Sequence 20 BP; 6 A; 3 C; 4 G; 7 T; 0 U; 0 Other;
                                           .
7
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Cote G,
             Score 15.8; DB 1;
Pred. No. 2e+02;
          . Match 0.4%; Score 15.8; D
Local Similarity 89.5%; Pred. No. 2e+0
les 17; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Falardeau P,
                                                                          2916 AAGGAACCCATGACAAAGA 2934
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Claim 18; Page 28; 41pp; English.
                                                                                                        19 AAGAAACCCAAGACAAAGA 1
                                                                                                                                                                                         BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  or progression of the disease
                                                                                                                                                                                                                                                                                      PCR primer for marker D6S967.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       98WO-CA000924.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      97CA-02217097.
                                                                                                                                                                                      AAX36672 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Morissette J,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 1999-263704/22.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (UYLA-) UNIV LAVAL
                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                      WO9916899-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    30-SEP-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      29-SEP-1998;
                                                                                                                                                                                                                                                      13-JUL-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      08-APR-1999.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Raymond V,
                                                                                                                                                                                                                                                                                                                                                                                      Synthetic
                                                                                                                                                                                                                      AAX36672;
             Query Match
                                          Matches
                                                                                                                                                         RESULT 155
```

Oligonucleotide for treating diseases associated with human tumor necrosis factor-alpha (TNF-alpha) such as, diabetes and rheumatoid arthritis, comprises nucleotide sequence complementary to intron of nucleic acid encoding TNF-alpha.

Example 22; Page 104; 283pp; English.

Butler MM, Shanahan WJ;

(ISIS-) ISIS PHARM INC. Baker BF, Bennett CF, WPI; 2000-303808/26.

18-MAY-1999;

05-OCT-1998;

99WO-US023205. 98US-00166186. 99US-00313932.

05-0CT-1999;

13-APR-2000.

```
ö
                                                                                                                                                                                                                                                                                                                                                                                           oligonuclectides optionally have a phosphorothicate backbone, and may also optionally contain at least one 2'-O-methoxyethyl modification. The oligonuclectides are useful for modulating the expression of human TNFalpha in cells and tissues, reducing a human cell inflammatory response, reducing the blood glucose level in a human and treating a human having a disease or condition associated with TNFalpha. Examples of diseases associated with TNFalpha include diabetes, inflammatory bowel
This sequence represents an antisense oligonucleotide sequence which targets a region of the human tumour necrosis factor alpha (TNFalpha) uncleotide sequence. The Alpha is an important cytckine that plays a role in host defence. It is produced mainly in macrophages and monocytes in response to infection, invasion, injury or inflammation. Overexpression of TNFalpha can result in disease states, particularly in infectious, inflammatory and autoimmune diseases. The invention relates to antisense oligonucleotides, such as that represented by the present sequence which are capable of modulating the TNFalpha gene expression. The
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       infectious disease, hepatitis, atopic dermatitis or allograft rejection.
The antisense oligonucleotides are also useful for modulating the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   disease, multiple sclerosis, pancreatitis, rheumatoid arthritis,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 function of a selected nucleic acid sequence in adipose tissue
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.4%; Score 15.8; DB 1; Length 20; 19.5%; Pred. No. 2e+02; ve 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Sequence 20 BP; 7 A; 4 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        3414 TTTCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              TCTCAAGGAAGTCTGGAAA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     83.58;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Best Local
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Matches
```

RESULT 157 AAF32850/c

8 셤

ö

Gaps

ö

DB 1; Length 20;

Score 15.8; DB 1; Length 20 Pred. No. 2e+02; 0; Mismatches 2; Indels

0.4%;

Query Match
Best Local Similarity 89.5
Matches 17; Conservative

3735 GGACCTATGAATCTATTTA 3753

ઠે

GCAACTGTGAATCTATTTA 19

-

RESULT 156

```
Novel antisense compound 8-30 nucleobases in length targeted to a nucleic acid molecule encoding human mdm-2 useful for modulating the expression of human mdm-2 and reducing hyperproliferation of human cells.
                                                                                                                                                                                                                                                                            The present invention relates to an antisense compound 8-30 nucleobases in length targeted to nucleobases 1-308 of the 5' untranslated region, 1776-1806 of the translation termination codon region or 1818-2370 of the 3' untranslated region of a nucleic acid molecule encoding human mdm-2. The invention is useful for reducing hyperproliferation of human cells, modulating the expression of mdm2 in human cells or tissues or in vitro. The hyperproliferative disorder includes cancer or psoriasis
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     /noce = "OTHER= All phosphorothioate linkages, additionally bases 1-6 and bases 15-20 are 2'-0-methoxyethyl bases, and bases 7-14 are deoxynucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Human, mdm2; hyperproliferative disorder; cancer; psoriasis; atherosclerosis; tumour; cytostatic; anti psoriatic; anti arteriosclerotic; vasotropic; antisense; phosphorothioate; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match 0.4%; Score 15.8; DB 1; Length 20; Best Local Similarity 89.5%; Pred. No. 2e+02; Matches 17; Conservative 0; Mismatches 2; Indels
                                                                                        Cowsert LM;
                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 0 A; 4 C; 4 G; 12 T; 0 U; 0 Other;
                                                                                          Graham MJ, Monia BP,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Human mdm2 antisense oligonucleotide 31768.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 1. .20
/*tag= a
/mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         2916 AAGGAACCCATGACAAAGA 2934
                                                                                                                                                                                                                                              Example 9; Col 31; 77pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             П
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ВР
                98US-00048810.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             19 AAGAAACCCAAGACAAAGA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  02-JAN-2001; 2001US-00752983
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        98US-00048810
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        99US-00280805
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             426/c
AAS29426 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (first entry)
                                                    (ISIS-) ISIS PHARM INC.
                                                                                          Nero P,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              MIRAGLIA L J.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  MONIA B P.
COWSERT L M.
                                                                                                                                WPI; 2001-190948/19.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  NERO P.
GRAHAM M J.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        US2001016575-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Key
modified base
                26-MAR-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      26-MAR-1998;
26-MAR-1999;
                                                                                            Miraglia LJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       21-NOV-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              23-AUG-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAS29426;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (MIRA/) I
(NERO/) I
(GRAH/) (
(MONI/) I
(COWS/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 159
셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ሯ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Novel compound for diagnosing, preventing and treating immune disorders, comprising an oligonucleotide that specifically hybridizes with a nucleic acid sequence encoding B7 protein.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 The present invention provides sequences of antisense oligonucleotides targeted at the murine and human B7-1 and B7-2 coding and mRNA sequences. The antisense sequences have phosphorothicate backbones and some nucleotides are 2'-methoxyethoxy residues. The sequences can be used in the treatment of inflammatory and autoimmune disorders, including asthma, juvenile diabetes mellitus, myasthenia gravis, Graves' disease, rheumatoid arthritis, allograft rejection, inflammatory bowel disease, multiple sclerosis, psoriasis, systemic lupus erythematosus, contact dermatitis, rhinitis, allergies and cancer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             .
0
                                                                                                                                                  Human, mouse, B7-1; B7-2; antisense; PCR primer; inflammation; autoimmune disorder; phosphorothioate backbone; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Antisense; mdm2; hyperproliferation; cancer; psoriasis; ss.
                                                                                                              Human B7-1 mRNA antisense oligonucleotide SEQ ID NO: 47
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 7 A; 4 C; 8 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human mdm2 phosphorothioate oligonucleotide #185.
                                                                                                                                                                                                                                                                                                                                                                                                                                            Karras JG;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Example 1; Page 47; 162pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 3766 TGGCTGGGATCCCTCCCT 3784
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20 rederedentecereter 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ВР
                                                                                                                                                                                                                                                                                                                            25-MAY-2000; 2000WO-US014471.
                                                                                                                                                                                                                                                                                                                                                                  99US-00326186.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            99US-00280805.
AAF32850 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AAF80811 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                                          (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                            Bennett CF, Vickers TA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                      (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2001-049991/06.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Local Similarity
                                                                                                                                                                                                                                                  WO200074687-A1
                                                                                                                                                                                                              Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          26-MAR-1999;
                                                                                                                                                                                                                                                                                                                                                                  04-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    US6184212-B1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              02-MAY-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        06-FEB-2001
                                                                                                                                                                                                                                                                                      14-DEC-2000
                                                                          23-MAR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
                                    AAF32850
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAF80811
```

Matches

ò 원 ö

Gaps

Monia BP, Cowsert LM;

Graham MJ,

Nero P,

```
Miraglia LJ,
```

```
length targeted to the 5 untranslated region, translation termination codon region, 3' untranslated region, coding region or translation start site of a nucleic acid encoding human mada, where the antisense compound modulates the expression of human mada. They may be used for treating an animal having a disease or condition associated with amplification of mada encoding overexpression of human mada. They may be used for treating an animal having a disease or condition associated with amplification of mada gene or overexpression of mada e.g. a hyperproliferative disorder such as cancer (blood, brain, breast, lung, or a soft tissue cancer) and psoriasis, fibrosis, atherosclerosis or restenosis, tumours, colorectal carcinoma and chronic myelogenous leukemia. The antisense compound may be administered with a chemotherapeutic agent to overcome drug resistance.
An antisense compound, useful for treating e.g. cancer, comprises nucleobases targeted a region (e.g. translation termination codon region) of a nucleic acid encoding human mdm2.
                                                                                                                                                                                                                                                                                                                                                      present invention relates to antisense compounds, 8-30 nucleobases in
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      The antisense compound reduces hyperproliferation of human cells. The method, which involves the use of the antisense compound, is also useful for detecting the role of mdm2 expression in various cell functions and physiological processes and useful in both clinical research and diagnostic tools. AAS29242-AAS29507 represent the human mdm2 antisense
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 20 BP; 0 A; 4 C; 4 G; 12 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            oligonucleotides of the present invention
                                                                                                                                                                                                                                    Example 9; Page 17; 81pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.4%;
```

```
ô
                            Gaps
                            ;
0
Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                     2916 AAGGAACCCATGACAAAGA 2934
              Local Similarity 89.5
ses 17; Conservative
  Query Match
                            Matches
                                                     ò
```

AAGAAACCCAAGACAAAGA 1

19

g

ABN79736 standard; DNA; 20 BP 29-JUL-2002 (first entry) ABN79736; RESULT 160 ABN79736/ 

Human Fas target oligonucleotide #51

Human; immunosuppressive; antiinflammatory; hepatotropic; cytostatic; vasotropic; hepatitis; cancer; allograft rejection; ds; Fas.

JS2002004490-A1. Homo sapiens.

10-JAN-2002.

09-MAR-2001; 2001US-00802669

12-APR-1999; 99US-00290640. 18-SEP-2000; 2000US-00665615.

(DEAN/) DEAN N M. (MARC/) MARCUSSON E G. (WYAT/) WYATT J. (ZHAN/) ZHANG H.

Zhang H; Wyatt J, Dean NM, Marcusson EG,

WPI; 2002-204886/26

```
ö
                 Novel antisense compound targeted to nucleic acid encoding Fas, Fas ligand or Fas associated protein-1 is useful for inhibiting expression of Fas, Fas ligand, or Fap-1 in cells or tissues, and for treating
                                                                                                                                                                            This invention relates to an antisense compound encoding Fas, Fas ligand, or Fas associated protein-1 (Fap-1). The inhibition of Fas mediated signalling is thought to be immunosuppressive, antiinflammatory, hepatotropic, cytostatic and vasotropic. Antisense oligonucleotides were designed to target human Fas. Oligonucleotides were synthesised as chimeric oligonucleotides and are useful for treating an animal having an autoimmune or inflammatory disease e.g., hepatitis, cancer, a condition associated with apoptosis, allograft rejection, or ischemia reperfusion injury. Optionally, the above mentioned conditions are prevented by contacting the allograft with the antisense oligonucleotide. The
                                                                                                                                                                                                                                                                                                                                                                                                         oligonucleotides are used in diagnostics, therapeutics, prophylaxis and as research reagents and in kits. The oligonucleotides are also useful for research purposes. The present nucleotide sequence is related to human Fas
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 20 BP; 9 A; 1 C; 0 G; 10 T; 0 U; 0 Other;
                                                                                                                                    Example 18; Page 24; 84pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        861 TTAAGAAATAATTTTGATA 879
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20 rradgaaaraararrara 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Best Local Similarity 89.5
Matches 17; Conservative
                                                                                   hepatitis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  d
```

Tumour necrosis factor alpha antisense oligonucleotide #370. (first entry) 05-AUG-2003 THE STATE OF THE S

ACD05367 standard; DNA; 20 BP.

RESULT 161 ACD05367 ACD05367;

Tumour necrosis factor alpha; TNF-alpha; antiinflammatory; antirheumatic; antiarthritic; antidiabetic; dermatological; hepatotropic; antiasthmatic; inflammatory bowel disease; Crohn's disease; colitis; rheumatory arthritis; diabetes; pancreatitis; multiple sclerosis; atopic dermatitis; asthma; hepatitis; antisense technology; ss

Synthetic.

US2003022848-A1. 30-JAN-2003.

98US-00166186. 99US-00313932. 02-APR-2001; 2001US-00824322 05-OCT-1998; 18-MAY-1999;

BUTLER M M. SHANAHAN W R. BAKER B F. BENNETT C F. (BENN/) 1 (BUTL/) I Shanahan WR; Butler MM, Bennett CF, Baker BF,

WPI; 2003-447433/42.

Treating inflammatory disorders such as inflammatory bowel disease, Crohn's disease or rheumatoid arthritis, in a subject, by administering

sequence

```
The invention comprises antisense oligonucleotides which are targeted to the human mdm2 gene. The antisense oligonucleotides of the invention are useful for reducing hyperproliferation of human cells. The antisense oligonucleotides are also useful for treating: hyperproliferative disorders (e.g. cancer), psoriasis, fibrosis, atheroselerosis, or restenosis. The antisense oligonucleotides are also useful for modulating
                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 2003-577263/54.
                                                                                                                                                                                                                                                                                                                                                                                            Nero
                                                                                                                                           Local Similarity
                                                                                                        alpha (TNF-alpha)
                                                                                                                                                                                                                                                                                                                       WO2003048315-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                     mdm2 expression
                                                                                                                                                                                                                                                                                                                                                                                             Miraglia LJ,
                                                                                                                                                                                                                                                                                                           Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                    Manoharan M;
                                                                                                                                                                                                                                                                                                                                      12-JUN-2003
                                                                                                                                                  17;
                                                                                                                                                                                                                              ADD21622;
                                                                                                                                    Query Match
                                                                                                                                                  Matches
                                                                                                                                                                                                 ò
                                                                                                                                                                           임
```

ö The invention describes a method of treating an inflammatory disorder in an individual, comprising administering to the individual an oligonucleotide upto 30 nucleotides in length complementary to a nucleic acid molecule encoding human tumor necrosis factor (TMP)-alpha. The method is useful for treating an inflammatory disorder such as inflammatory bowel disease, Crohn's disease, colitis or rheumatoid arthritis, in an individual. The method is also useful for treating diabetes, pancreatitis, multiple sclerosis, atopic dermatitis, asthma, and hepatitis in an individual. This sequence represents an antisense oligonucleotide which inhibits expression of human tumor necrosis factor Novel antisense compound targeted to 5' untranslated region, coding region, or intron:exon junction of nucleic acid molecule encoding mdm2, useful for treating e.g. cancer, psoriasis or restenosis by inhibiting oligonucleotide used to modulate expression of tumour necrosis factor Gaps Chiang MY; ô antisense oligonucleotide; human; mdm2; hyperproliferation; hyperproliferative disorder; cancer; psoriasis; fibrosis; atherosclerosis; restenosis; apoptosis modulation; p21; ss; 2'-methoxyethoxy-residue; phosphorothioate backbone. Score 15.8; DB 1; Length 20; Pred. No. 2e+02; 2; Indels Monia BP, Koller E, Sequence 20 BP; 7 A; 4 C; 5 G; 4 T; 0 U; 0 Other; 2e+02; 0; Mismatches Human mdm2 antisense oligonucleotide #185. Example 9; SEQ ID NO 187; 289pp; English. Example 24; Page 39; 142pp; English Graham MJ, 3414 TTTCAAGGAAGTATGGAAA 3432 BP 0.4%; 02-DEC-2002; 2002WO-US038281. 04-DEC-2001; 2001US-00005344 ADD21622 standard; DNA; 20 15-JAN-2004 (first entry) Conservative PS, (ISIS-) ISIS PHARM INC

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
apoptosis, and for increasing expression of p21. The present DNA sequency represents a human mdm2 gene antisense oligonucleotide of the invention. The present sequence contains 2'-methoxyethoxy-residues and has a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               disorder in an individual by topically applying an antisense compound targeted to a nucleic acid molecule encoding a human B7 protein. The invention is for treating an inflammatory skin disorder in individual. The skin disorder is psoriasis, contact dermatitis, atopic dermatitis, seborrheic dermatitis, nummular dermatitis, generalised exfoliative dermatitis or eczema. The invention effectively modulates critical costinulatory molecules such as the B7 protein. The present sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Treating an inflammatory skin disorder such as psoriasis comprises topically applying an antisense compound targeted to the nucleic acid encoding human B7 protein.
                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                  ss; human; B7-1; inflammatory skin disorder; antisense; psoriasis; conteact dermatitis; atopic dermatitis; seborrheic dermatitis; nummular dermatitis; generalised exfoliative dermatitis; eczema; critical costimulatory molecule.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  invention relates to a method of treating an inflammatory skin
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ..
0
                                                                                                                                              ö
                                                                                                           Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                      Human B7-1 mRNA targeted oligonucleotide SEQ ID 47.
                                                                                 Sequence 20 BP; 0 A; 4 C; 4 G; 12 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 20 BP; 7 A; 4 C; 8 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              represents a human B7-1 targeted oligonucleotide.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Example 1; SEQ ID NO 47; 88pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Karras JG;
                                                                                                                                                                             2916 AAGGAACCCATGACAAAGA 2934
                                                                                                                                                                                                             AAGAAACCCAAGACAAAGA 1
                                                                                                                                                                                                                                                                                           BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                09-MAY-2001; 2001US-00851871.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              96US-00777266
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           04-JUN-1999; 99US-00326186.
25-MAY-2000; 2000WO-US014471.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match

Best Local Similarity 89.5%;
Matches 17; Conservative
                                                                                                                  0.4%;
                                                                                                                                                                                                                                                                            785/c
ADE27785 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                         (first entry)
                                                 phosphorothioate backbone
                                                                                                      Query Match
Best Local Similarity 89.5
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Bennett CF, Vickers TA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           BENNETT C F.
VICKERS T A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2003-863863/80.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          KARRAS J G.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   US2003176374-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               31-DEC-1996;
                                                                                                                                                                                                                                                                                                                                                         29-JAN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  18-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Synthetic
                                                                                                                                                                                                             19
                                                                                                                                                                                                                                                                                                                          ADE27785;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (VICK/)
(KARR/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (BENN/)
                                                                                                                                                                                                                                                             RESULT 163
                                                                                                                                                                                                                                                                             ADE27785
                                                                                                                                                                                                                                                                                                           8886666
                                                                                                                                                                                                             g
                                                                                                                                                                                 ਨੇ
```

~

Human, antisense, lung dysfunction, nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiathmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; 3766 TGGCTGGGATCCCTCCCCT 3784 Human oligonucleotide sequence ABZ85958 standard; DNA; 20 BP. rederedecarecterer 17-OCT-2003 (first entry) WPI; 2003-229219/22. WO200285308-A2 Homo sapiens. 31-OCT-2002 Nyce JW, I Miller S, ubiquinone 20 ABZ85958; RESULT 164 ABZ85958 g ŝ

lung inflammation; respiratory disease; ds Katz E, Li Y, Sandrasagra A, Ka Tang L, Shahabuddin S; 23-APR-2002; 2002WO-US013135. 24-APR-2001; 2001US-0286137P. (EPIG-) EPIGENESIS PHARM INC

Pabalan J, Aguilar D;

Pharmaceutical composition for treating ailments associated with impaired or respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid

Claim 15; SEQ ID NO 1200; 872pp; English.

The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation coodon, coding regalon, 5' or 3' end ganomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an entilnflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallargic, antiasthmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a use in antisense gene therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of an antiinflammatory steroid in a subject, for reducing levels of condition, and of, or reducing sensitivity to adenosine, reducing levels of using surfactant in a subject or treating bronchoconstriction, lung surfactant in a subject or a respiratory disease or condition. Note: The sequence date for this patent is not represented in the printed specification, but was obtained in electronic format directly from WIPO at ftp.wipo.int/pub/published\_pct\_sequences

Sequence 20 BP; 5 A; 1 C; 7 G; 7 T; 0 U; 0 Other;

Score 15.8; DB 1; Length 20; Pred. No. 2e+02; 0; Mismatches 2; Indels 89.5%; Query Match 0.4 Best Local Similarity 89.5 Matches 17; Conservative

ö

Gaps

. 0

2092 TAGTATCTGTTGTAGCAGT 2110 13 1 TAGTATCTGTTGGAGAGT 셤 ò

ABZ90805 standard; DNA; 20 BP ABZ90805

RESULT 165

ABZ90805;

(first entry) 17-OCT-2003 Human oligonucleotide sequence

Human, antisense; lung dysfunction; nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiinflammatiory steroid; ubiquinone; antiinflammatory; antiallergic; antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodiation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds.

Homo sapiens

WO200285308-A2. 

31-OCT-2002.

23-APR-2002; 2002WO-US013135.

24-APR-2001; 2001US-0286137P.

(EPIG-) EPIGENESIS PHARM INC.

Pabalan J, Aguilar D; M, Katz Li Y, Sandrasagra A, K Tang L, Shahabuddin S; Miller S, Nyce JW

WPI; 2003-229219/22.

Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or ubiquinone

Disclosure; SEQ ID NO 6047; 872pp; English

lung inflammation, lung allergies, or a respiratory disease or condition. Note: The sequence data for this patent is not represented in the printed specification, but was obtained in electronic format directly from WIPO The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonuclectide antieense to the initiation codon, coding region, 5' or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or ansal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiasthmatic, hypotensive, ung surfactant a subject's tissue, or treating bronchoconstriction, immunosuppressive, and cytostatic activity. The composition may have a use in antisense gene therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of an antiinflammatory steroid in a subject, for reducing or depleting levels of, or reducing sensitivity to adenosine, reducing levels of adenosine receptor, producing bronchodilation, increasing levels of ubiquinone or at ftp.wipo.int/pub/published\_pct\_sequences

Sequence 20 BP; 1 A; 6 C; 1 G; 12 T; 0 U; 0 Other;

0; Gaps 0.4%; Score 15.8; DB 1; Length 20; 89.5%; Pred. No. 2e+02; 2; Indels 2; Indels Query Match 0.4 Best Local Similarity 89.5 Matches 17; Conservative

```
TGTTCACTTTTTCCTTTTT 3099
                 refrcacirriricercre 19
3081
                 н
ઠ
                 g
```

ABZ93212 standard, DNA; 20 BP

ABZ93212;

17-OCT-2003 (first entry)

Human oligonucleotide sequence

Human, antisense; lung dysfunction, nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; adenosine receptor; bronchodilation; bronch lung inflammation; respiratory disease; ds.

Homo sapiens.

WO200285308-A2

31-OCT-2002.

23-APR-2002; 2002WO-US013135.

24-APR-2001; 2001US-0286137P.

(EPIG-) EPIGENESIS PHARM INC.

Katz E, Pabalan J, Aguilar D; Li Y, Sandrasagra A, Ka Tang L, Shahabuddin S; Nyce JW, | Miller S,

WPI; 2003-229219/22.

Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or ubiquinone

Disclosure; SEQ ID NO 8454; 872pp; English.

The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation coodon, coding regaion, 5' or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallargic, antiathmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a use in antisense gene therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of an expectivity to adenosine. reducing levels of adenosine receptor, producing sensitivity to adenosine, reducing levels of adenosine receptor, producing sensitivity to adenosine, reducing levels of adenosine receptor, producing sensitivity to adenosine, reducing levels of adenosine lung surfactant in a subject's tissue, or treating bronchoconstriction, lung inflammation, lung allergies, or a respiratory disease or condition. Note: The sequence date for this patent is not represented in the printed appearance of the printed and the printed printed and the printed at ftp.wipo.int/pub/published\_pct\_sequences ABZ93212
XXD ABZ9222
XXD

Sequence 20 BP; 12 A; 4 C; 4 G; 0 T; 0 U; 0 Other;

```
Gaps
                               ;
 Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
     0.4%;
Query Match
Best Local Similarity 89.5
Matches 17; Conservative
```

ö

2974 CAGAGAAACAGAAAGAGA 2992 20 2 CAGAGCAAACAGAAAAAGA 셤 ò

ABZ98920

ВР ABZ98920 standard; DNA; 20

(first entry) 17-0CT-2003 ABZ98920;

Human PDE4A oligonucleotide sequence.

antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; tespiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds. Human; antisense; lung dysfunction; nasal airway dysfunction;

Homo sapiens.

WO200285308-A2.

31-OCT-2002.

23-APR-2002; 2002WO-US013135.

24-APR-2001; 2001US-0286137P.

(EPIG-) EPIGENESIS PHARM INC.

ä Pabalan J, Aguilar Katz E, ŝ Li Y, Sandrasagra A, Tang L, Shahabuddin Nyce JW, I Miller S,

WPI; 2003-229219/22.

Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or ubiquinone

Disclosure; SEQ ID NO 14162; 872pp; English.

The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation codon, coding region, 5' or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiasthmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a cue in antisense gene therapy. The composition is useful for treating or preventing the prophylactic or therapeutic respiratory effect of an oriniflammatory steroid in a subject, for reducing levels of continion or infinitammatory steroid in a subject, for reducing levels of adenosine receptor, producing bronchodilation, increasing levels of adenosine receptor, producing bronchodilation, increasing levels of ubiquinone or lung inflammation, lung allergies, or a respiratory disease or condition, lung inflammation, lung allergies, or a respiratory disease or condition. Note: The sequence data for this patent is not represented in the printed specification, but was obtained in electronic format directly from WIPO at ftp.wipo.int/pub/published\_pct\_sequences 

Sequence 20 BP; 3 A; 6 C; 8 G; 3 T; 0 U; 0 Other;

Gaps ö 0.4%; Score 15.8; DB 1; Length 20; 89.5%; Pred. No. 2e+02; tive 0; Mismatches 2; Indel8 Query Match 0.4 Best Local Similarity 89.5 Matches 17; Conservative

Toxicologically relevant gene; toxicological response; PCR primer; ss.

16-AUG-2002; 2002WO-US026514.

WO2003016500-A2 27-FEB-2003

Rattus sp. Synthetic.

Toxicologically relevant rat PCR primer #1056.

14-MAY-2003 (first entry)

ABZ83897;

8

Page

2707 GAGGCATTTCTTGCCCAGC 2725

ò 셤

1 GAGGCATTTCTTGTCCACC 19

ABZ83897 standard; DNA; 20 BP

ABZ83897,

```
at ftp.wipo.int/pub/published_pct_sequences
       ~
                                                                                                             23-APR-2002; 2002WO-US013135.
      20 CCGCTGGAGCTGTACCGCC
                                                                                                                     24-APR-2001; 2001US-0286137P.
                                                                                                                             (EPIG-) EPIGENESIS PHARM INC.
                          ABZ97844 standard; DNA; 20
                                          17-OCT-2003 (first entry)
                                                                                                                                                WPI; 2003-229219/22.
                                                                                              WO200285308-A2
                                                                                      Homo sapiens.
                                                                                                     31-OCT-2002.
                                                                                                                                                                     ubiquinone
                                                                                                                                         Miller S,
                                  ABZ97844;
                                                                                                                                     Š
                   RESULT 168
                                                                                                                                      Nyce
ઠે
      셤
```

```
The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonuclectide antisense to the initiation codon, coding region, 5 or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubjquinone. A composition of the invention has antiinflammatory, antiallergic, antiasthmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a use in antisense gene therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of an antiinflammatory steroid in a subject, for reducing levels of, or reducing sensitivity to adenosine, reducing levels of adenosine or processing producing producing producing increasing levels of uniquinone or processive.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                lung surfactant in a subject's tissue, or treating bronchoconstriction, ung inflammation, lung allergies, or a respiratory disease or condition. Note: The sequence data for this patent is not represented in the printed specification, but was obtained in electronic format directly from WIPO
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human; antisense; lung dysfunction; nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Katz E, Pabalan J, Aguilar D;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Disclosure; SEQ ID NO 13086; 872pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Human eotaxin oligonucleotide sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Li Y, Sandrasagra A, Ka
Tang L, Shahabuddin S;
2508 CAGGIGGAGCTGIACCGCC 2526
```

```
reference gene expression profile indicative of toxicity, and so determining the presence of a toxic response to the agent. Also determining the presence of a toxic response to the agent. Also determining the presence of a toxic response to the agent. Also from the genes corresponding to the partial sequences given in AB282842 to AB284764, or their fragments of at least 20 nucleotides, or homologues of and (2) determining if a gene putatively identified to be a toxic response gene plays a role on toxic response pathways by determining the expression profile of the gene after exposure of cells or a human subject to a known toxic pharmaceutical or industrial agent, comprising; (a) exposing cells to an agent or isolating cells from a human subject who was exposed to an agent or isolating the test gene expression profile for a putatively identified toxic response gene after exposure to a known coxic pharmaceutical or industrial agent; and (c) comparing the test profile to the expression profile of a gene with a similar function or comparing the test profile to the expression profile of a gene with a similar function or profile to the test profile to the expression profile of a gene with a similar function or profile to the expression profile of a gene with a similar function or comparing the test profile to the expression profile of a gene with a similar function or predicting and determining toxicological responses on a cellular, organ or system level. The arrays comprising the human genes are useful for toxicological screening of drugs, pharmaceutical compounds and chemicals
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Determining a toxicological response to an agent, useful for screening of drugs, comprises comparing the expression profile of one or more human toxic response genes to a reference gene expression profile indicative of
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Kier LD, Schmeiser K;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             The present invention describes a method (M1) for determining a toxicological response to an agent, which comprises comparing the expression profile of one or more human toxic response genes to a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match 0.4%; Score 15.8; DB 1; Length 20; Best Local Similarity 89.5%; Pred. No. 2e+02; Matches 17; Conservative 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 20 BP; 3 A; 7 C; 3 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Neft RE, Dunn RT, Adkins K, Pickett GG,
Alen P;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (PHAS-) PHASE-1 MOLECULAR TOXICOLOGY INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Claim 1; Page 322; 455pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  2653 GATGCAATTGGCAGGAAGC 2671
                                                                                                                                                                                                                                                                                                                                                                                                                                                                    16-AUG-2001; 2001US-0313080P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2003-268322/26.
```

λō.

ö

Gaps

; 0

Score 15.8; DB 1; Length 20; Pred. No. 2e+02; 0; Mismatches 2; Indels

Sequence 20 BP; 3 A; 7 C; 4 G; 6 T; 0 U; 0 Other;

0.4%;

Query Match
Best Local Similarity 89.5
Matches 17; Conservative

10001863-3.81.rng

ss; retroelement; detection; retrovirus; plant.

WO2003050259-A2.

Synthetic.

.9-JUN-2003

Plant retroelement associated sequence #25

(first entry)

06-MAY-2004

ADJ81464;

```
The present invention describes a DNA molecule derived from a prokaryotic cell, where at least one of the DNA regions encoding NNB (where N = asparagine, X = any amino acid other than proline, and B = serine or threonine) has been modified so that no N-glycosylation occurs during the expression in a eukaryotic cell. Also described: (1) a fused DNA molecule, where a DNA encoding a signal sequence has been ligated to the N-terminal end of the modified DNA molecule as described above so that it may be expressed as a fusion protein; (2) a recombinant virus integrated with the DNA molecule or the fused DNA molecule described above; (3) producing a modified or fusion protein by using the recombinant virus described above, to express a protein encoded by the modified DNA molecule or the fused DNA molecule in a eukaryotic cell; and (4) a vaccine comprising the recombinant virus. The DNA molecule has virucide a vaccine for treating viral infections. The DNA molecule is useful for producing a vaccine for treating viral infections. The present sequence is used in the exemplification of the present invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DNA molecule derived from a prokaryotic cell, useful for producing a vaccine for treating viral infections comprises at least one modified DNA regions encoding NXB so that no N-glycosylation occurs during expression.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           DNA molecule, prokaryotic cell; eukaryotic cell, virucide, vaccine, immunostimulant, viral infection; PCR primer; linker; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                    M. gallisepticum mgc3 gene BKR region mutagenic PCR primer SEQ:36.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ch 0.4%; Score 15.8; DB 1; Length 20; 1 Similarity 89.5%; Pred. No. 2e+02; 17; Conservative 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 20 BP; 9 A; 2 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Tsuzaki Y;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Example 2; Page 49; 70pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Dorsey KM,
                                                                                                                                                                                                                                      ΒÞ
GGTGCAATTGGCAAGAAGC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                11-JUL-2002; 2002EP-00254879.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        11-JUL-2001; 2001US-00901572.
25-APR-2002; 2002US-00131591.
                                                                                                                                                                                                                                          20
                                                                                                                                                                                                                                                                                                                                                                                            (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Mycoplasma gallisepticum.
                                                                                                                                                                                                                                      ACF03393 standard; DNA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2003-373746/36.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Saito S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (JAPG ) ZEON CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             EP1275716-A2
                                                                                                                                                                                                                                                                                                                                                                                            11-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         15-JAN-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Okuda T,
                                           13
                                                                                                                                                                                                                                                                                                                 ACF03393;
                                                                                                                                                          ACFO3393/C
ID ACFO3393/C
XX
XX
XX
XX
DD M. gal
XXX
XX
DD M. gal
XXX
MYCOPI
OS MYCOPI
O
                                               硆
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human, antisense; bronchoconstriction, allergy; hyposecretion; pain; respiratory tract inflammation; adenosine sensitivity; lung; cancer; surfactant depletion; antiallergic; antiinflammatory; antiasthmatic; analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic finititis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    The invention relates to novel isolated retroelement comprising at least 90% identity to a 14028 base pair sequence (S1), given in the specification or to nucleotides 1-1747, 12220-13864, 1-385, 1-40, 1708-1747, 1893-3575, 3576-4555, 4602-6314, 6315-7655, 8745-10600, 8745-10673 or 8745-10728 of this sequence, or its complement. The nucleic acid may be useful in detecting infections, such as those caused by retroviruses. This sequence corresponds to a retroelement associated sequence.
                                                                                                                                                                                                                                                                                                                                                                                          New isolated nucleic acids related to retrovirus elements from
Arabidopsis thaliana, useful for detecting infections, such as those
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%; Score 15.8; DB 1; Length 20;
89.5%; Pred. No. 2e+02;
ive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 5 A; 5 C; 4 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    H93087-derived oligonucleotide SEQ ID 6047.
                                                                                                                                                                                                                                                                                                                                                                                                                                                         Disclosure, SEQ ID NO 63; 81pp; English
                                                                                                                                                                                                                                                                                             (IOWA ) UNIV IOWA STATE RES FOUND INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2465 CITCATIGICCIGCAGAAG 2483
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      20 cargarrerecrecadade 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ABD27035 standard; DNA; 20 BP
                                                                                                                                                                                                                          10-DEC-2002; 2002WO-US039397.
                                                                                                                                                                                                                                                           10-DEC-2001; 2001US-0339060P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Local Similarity 89.5
les 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                            caused by retroviruses.
                                                                                                                                                                                                                                                                                                                           Wright DA, Voytas DF;
                                                                                                                                                                                                                                                                                                                                                          WPI; 2003-532907/50.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WO200285309-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      29-JUL-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  31-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ABD27035;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RESULT 172
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    셤
```

.; 0

Gaps

; 0

3702 TGTTTATTTTTCAGAAC 3720

ò 원

Best Loca Matches

rerrrarrererreadade 1

13

ADJ81464 standard; DNA; 20 BP.

ADJ81464/c ID ADJ814 XX

RESULT 171

17; Miller S, Nyce JW, Query Match Matches q 8

```
This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating bronchoconstriction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung dysfunction or cancer and can be anti-sense to the corresponding mkNA. The invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antiinflammatory, antiasthmatic,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            pulmonary obstruction, and/or bronchoconstriction and/or lung inflammation, allergies and/or surfactant hypoproduction are associated with a disease or condition such as pulmonary vasoconstriction, inflammation, allergies, asthem, impeded respiration, respiratory distress syndrome, pain, cystic fibrosis, allergic rhinitis, pulmonary hypertension, emphysema, chronic obstructive pulmonary disease, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of the oligonucleorides into products that free adenosine into the system
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ung, brain, heart, kidney, etc, tissue environment and thereby, to
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 analgesic, hypotensive, immunosuppressive and cytostatic activity, is a beta-adrenergic agonist. The composition is useful for preventing or treating a respiratory, lung or malignant disease. The administered composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypeptide present in the lungs. The
                                                                                                                                                                                                                                                                                                        t
                                                                                                                                                                                                                                                                          Pharmaceutical composition for treating asthma, has antisense oligonucleotide containing less percentage of adenosine, targeted truncleic acids associated with lung airway or lung dysfunction, and
                                                                                                                                                         Katz E, Pabalan J, Aguilar D;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 20 BP; 1 A; 6 C; 1 G; 12 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                 Claim 15; SEQ ID NO 6047; 763pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            prevent any unwanted effects due to it
                                                                                                                                                         Li Y, Sandrasagra A, Ka
Tang L, Shahabuddin S;
                 23-APR-2002; 2002WO-US013143.
                                                             24-APR-2001; 2001US-0286036P.
                                                                                                          (EPIG-) EPIGENESIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                  bronchodilating agent.
                                                                                                                                                                                                                           WPI; 2003-093058/08.
```

ö Gaps ö Score 15.8; DB 1; Length 20; Pred. No. 2e+02; 2; Indels 0; Mismatches 3081 TGTTCACTTTTTCCTTTTT 3099 0.4%; Conservative Local Similarity

1 IGTICACTITITICCTCT 19

Human PDE4A-derived oligonucleotide SEQ ID 14162.

Human, antisense, bronchoconstriction; allergy, hyposecretion, pain, respiratory tract inflammation, adenosine sensitivity, lung, cancer,

beta-adrenergic agonist, respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer. analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; surfactant depletion; antiallergic; antiinflammatory; antiasthmatic;

Homo sapiens.

WO200285309-A2

31-OCT-2002.

23-APR-2002; 2002WO-US013143.

24-APR-2001; 2001US-0286036P.

(EPIG-) EPIGENESIS PHARM INC.

Pabalan J, Aguilar D; Katz E, Li Y, Sandrasagra A, Ka Tang L, Shahabuddin S; Miller S, Nyce JW,

WPI; 2003-093058/08.

ដ Pharmaceutical composition for treating asthma, has antisense oligonucleotide containing less percentage of adenosine, targeted trucleic acids associated with lung airway or lung dysfunction, and bronchodilating agent.

Claim 15; SEQ ID NO 14162; 763pp; English.

This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating comprising oligonucleotides, effective for alleviating comprising oligonucleotides, effective for alleviating reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The coligonucleotides are derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung expression of a target polypeptide associated with lung airway or lung cyptention also describes a kit, that comprises: (a) a delivery device, in separate containers (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallargic, antiinflammatory, antiasthmatic, analgesic, hypotensive, immunosuppressive and cytostatic activity, is beta-adrenergic agonist. The composition is useful for preventing or treating a respiratory, lung or malignant disease. The administered composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to composition and/or bronchoconstriction and/or lung inflammation, allergies and/or surfactant hypoproduction are associated inflammation, allergies and/or surfactant hypoproduction are associated inflammation. inflammation, allergies, asthma, impeded respiration, respiratory distress syndrome, pain, cystic fibrosis, allergic rhinitis, pulmonary hypertension, emphysema, chronic obstructive pulmonary disease, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of oligonucleotides into products that free adenosine into the system

e.g., lung, brain, heart, kidney, etc, tissue environment and thereby, to prevent any unwanted effects due to it

Sequence 20 BP; 3 A; 6 C; 8 G; 3 T; 0 U; 0 Other;

Score 15.8; DB 1; Length 20; Pred. No. 2e+02; 0; Mismatches 2; Indels 2508 CAGGTGGAGCTGTACCGCC 2526 Query Match 0.4 Best Local Similarity 89.5 Matches 17; Conservative

89.58;

N

20 ccecrecaecreraccecc

g ઠ

ö

Gaps

Sequence 20 BP; 5 A; 1 C; 7 G; 7 T; 0 U; 0 Other;

```
analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer.
                                                                    Human, antisense; bronchoconstriction; allergy; hyposecretion; pain; respiratory tract inflammation; adenosine sensitivity; lung; cancer; surfactant depletion; antiallergic; antiinflammatory; antiasthmatic;
                                                    Human stanniocalcin-derived oligo SEQ ID 1200.
                                                                                                                                                                                                                                                               Sandrasagra A,
ABD22188 standard; DNA; 20 BP
                                                                                                                                                                                                           23-APR-2002; 2002WO-US013143.
                                                                                                                                                                                                                            24-APR-2001; 2001US-0286036P.
                                                                                                                                                                                                                                            EPIG-) EPIGENESIS PHARM INC
                                  29-JUL-2004 (first entry)
                                                                                                                                                                                                                                                                                         WPI; 2003-093058/08.
                                                                                                                                                                                                                                                               Li Y, San
Tang L,
                                                                                                                                                                      WO200285309-A2.
                                                                                                                                                      Homo sapiens
                                                                                                                                                                                        31-OCT-2002.
                                                                                                                                                                                                                                                                       Miller S,
                ABD22188;
                                                                                                                                                                                                                                                               Ę,
                                                                                                                                                                                                                                                                Nyce
```

to oligonuclectide containing less percentage of adenosine, targeted t nucleic acids associated with lung airway or lung dysfunction, and Pharmaceutical composition for treating asthma, has antisense bronchodilating agent.

Aguilar D;

Katz E, Pabalan J,

ŝ

Claim 15; SEQ ID NO 1200; 763pp; English

inflammation, allergies and/or surfactant hypoproduction are associated with a disease or condition such as pulmonary vasoconstriction, thilammation, allergies answer as pulmonary vasoconstriction, allergies asthma, impeded respiration, respiration, distress syndrome, pain, cystic fibrosis, allergic rhinitis, pulmonary hypertension, emphysema, chronic obstructive pulmonary disease, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of the oligonucleotides into products that free adenosine into the system e.g., lung, brain, heart, kidney, etc, tissue environment and thereby, to prevent any unwanted effects due to it comprising oligonucleotides, effective for alleviating bronchoconstriction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung dysfunction or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antiinflammatory, antiasthmatic, analgesic, hypotensive, immunosuppressive and cytostatic activity, is a beta-adrensful agonist. The composition is useful for preventing or treating a respiratory, lung or malignant disease. The administered composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypethide present in the lungs. The pulmonary obstruction, and/or bronchoconstriction and/or lung invention describes a novel composition (a) a first active agent,

```
This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating comprising oligonucleotides, effective for alleviating bronchoconstriction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, configonucleotides are derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung expression of a target polypeptide associated with lung airway or lung dysfunction or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antialflammatory, antiasthmatic, analgesic, hypotensive, immunosuppressive and cytostatic activity, is a conference respiratory, lung or malignant disease. The administered composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypeptide present in the lungs. The
                                    ö
                                                                                                                                                                                                                                                                                                                                                                               respiratory tract inflammation; adenosine sensitivity; lung; cancer; surfactant depletion; antiallergic; antiinflammatory; antiasthmatic; analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ဌ
                                    Gaps
                                                                                                                                                                                                                                                                                                                                                                     Human; antisense; bronchoconstriction; allergy; hyposecretion; pain;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Pharmaceutical composition for treating asthma, has antisense oligonucleotide containing less percentage of adenosine, targeted tnucleic acids associated with lung airway or lung dysfunction, and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ä
                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Pabalan J, Aguilar
DB 1; Length 20;
                                    2; Indels
                                                                                                                                                                                                                                                                                                                            Human eotaxin-derived oligonucleotide SEQ ID 13086
Score 15.8; DB 1 Pred. No. 2e+02;
                                      0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Claim 15; SEQ ID NO 13086; 763pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ы,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Katz
                                                                          2092 TAGTATCTGTTGTAGCAGT 2110
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ndrasagra A,
Shahabuddin
                                                                                                                13
                                                                                                                                                                                                             BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               23-APR-2002; 2002WO-US013143.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     24-APR-2001; 2001US-0286036P.
                                                                                                              1 ragrarcrerregadaadr
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (EPIG-) EPIGENESIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sandrasagra
                89.5%;
                                                                                                                                                                                                             ABD30875 standard; DNA; 20
                                                                                                                                                                                                                                                                                          (first entry)
                                        Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       bronchodilating agent.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Li Y, San
Tang L,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2003-093058/08.
                  Local Similarity
Les 17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WO200285309-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens.
                                                                                                                                                                                                                                                                                            29-JUL-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           31-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Miller S,
                                                                                                                                                                                                                                                      ABD30875;
Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Nyce JW,
                                                                                                                                                                           RESULT 175
                                        Matches
                                                                                                                                                                                                 ABD30875
                                                                                                                                                                                                                                   원
```

83

pulmonary obstruction, and/or bronchoconstriction and/or lung

```
with a disease or condition such as pulmonary vasoconstriction, inflammation, allergies, asthma, impeded respiration, respiratory distress syndrome, pain, cystic fibrosis, allergic rhinitis, pulmonary hypertension, emphysema, chronic obstructive pulmonary disease, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of the oligonucleotides into products that free adenosine into the system e.g., lung, brain, heart, kidney, etc, tissue environment and thereby, to prevent any unwanted effects due to it
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating bronchoconstriction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene encoding or regulating
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Human; antisense; bronchoconstriction; allergy; hyposecretion; pain; respiratory tract inflammation; adenosine sensitivity; lung; cancer; surfactant depletion; antiallergic; antiinflammatory; antiasthmatic; analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distresses syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; se; primer.
                       inflammation, allergies and/or surfactant hypoproduction are associated
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Pharmaceutical composition for treating asthma, has antisense oligonuclectide containing less percentage of adenosine, targeted to nucleic acids associated with lung airway or lung dysfunction, and
                                                                                                                                                                                                                                                                                                Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                               Sequence 20 BP; 3 A; 7 C; 4 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  H87536-derived oligonucleotide SEQ ID 8454.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Claim 15; SEQ ID NO 8454; 763pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Li Y, Sandrasagra A, Ka
Tang L, Shahabuddin S;
                                                                                                                                                                                                                                                                                                                                                                                    2707 GAGGCATTTCTTGCCCAGC 2725
                                                                                                                                                                                                                                                                                                                                                                                                                          1 GAGGCATTTCTTGTCCACC 19
                                                                                                                                                                                                                                                                                                       0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   23-APR-2002; 2002WO-US013143.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        24-APR-2001; 2001US-0286036P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (EPIG-) EPIGENESIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ABD29442 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          29-JUL-2004 (first entry)
                                                                                                                                                                                                                                                                                                                     Local Similarity 89.5
nes 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     bronchodilating agent.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WPI; 2003-093058/08.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WO200285309-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           31-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Nyce JW, I
Miller S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ABD29442;
                                                                                                                                                                                                                                                                                                     Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              tches
ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                    g
```

Katz E, Pabalan J, Aguilar D;

```
dysfunction or cancer and can be anti-sense to the corresponding arrays of the invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antiinflammatory, antiasthmatic, analgesic, hypotensive, immosuppressive and cytosteric activity, is a beta-adrenergic agonist. The composition is useful for preventing or treating a respiratory, lung or malignant disease. The administered composition comprises of oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or reduce the amount of target polypeptide present in the lungs. The reduce the amount of target polypeptide present in the lungs. The composition, allergies and/or surfactant hypoproduction are associated with a disease or condition such as pulmonary vasoconstriction, inflammation, allergies, asthma, impeded respiration, respiratory distress syndrome, pain, cystic fibrosis, allergic thintis, pulmonary hypertension, emphysema, chronic obstructive pulmonary disease, pulmonary corresponding to the included adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of the oligon pulmonary interpression, emphysema.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         e.g., lung, brain, heart, kidney, etc, tissue environment and thereby, to prevent any unwanted effects due to it
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               human, T-cell associated disease, Vbeta; autoimmune disease; degenerative nervous system disease; graft versus host disease; hypersensitivity disease; infectious disease; neoplastic disease; Addison's disease; atrophic gastritis; degenerative nervous system disease; multiple sclerosis; Alzheimer sdisease; hypersensitivity disease; type I hypersensitivity; Goodpasture's syndrome; type II hypersensitivity; Goodpasture's syndrome; type IV hypersensitivity; leprosy; infections disease; viral infection; HIV; fungal infection, Candida; parasitic infection; schistosome; filaria; bacterial infection; Mycobacterium; neoplastic disease; lymphoproliferative disease; leukaemia; lymphoma; cancer; brain cancer;
expression of a target polypeptide associated with lung airway or lung
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 12 A; 4 C; 4 G; 0 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Human Vbeta gene repeat sequence #192.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            2974 CAGAGAAACAGAAAGAGA 2992
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  2 CAGAGCAAACAGAAAAGA 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ADH70402 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              94US-00309335.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     99US-00263959.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     25-MAR-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ι.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  oreast cancer; ds
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (HOOD/) HOOD L E (ROWE/) ROWEN L.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  US2002150891-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                19-SEP-1994;
19-SEP-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     05-MAR-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             L7-OCT-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADH70402;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
Best Local S
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      RESULT 177
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ADH70402
  888888888888888888888888888888888
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  d
```

ö

ö

Rowen L;

Έ,

```
The invention relates to a kit for diagnosing and treating T-cell associated diseases which comprises a panel of nucleic acid primers specifically priming and allowing amplification of each Vbeta gene, VbetaRNA or cDNA. The kit is useful for diagnosing organ transplant respection and diagnosing and treating T-cell associated diseases.

The including autoimmune disease, hypersensitivity diseases, infectious diseases and neoplastic disease. Autoimmune diseases include Addison's disease.

The propriety of allowing and treative nervous system diseases include multiple actrophic gastritis. Degenerative nervous system diseases include multiple actrophic gastritis as contact with allergement in the sort as contact with allergement in the goodpasture's syndrome and Type IV hypersensitivities such as those caused by viruses such as those present in leprosy. Infectious diseases include viral infections caused by viruses such as HIV, fungal infections such as those caused by schildrand parterial infections such as those caused by childrand bacterial infections such as those caused by situation and bacterial infections such as those caused by such as leukaemias, lymphomas and cancers such as caused by hyperal and bacterial infections such as those caused by such as leukaemias, lymphomas and cancers such as caused by contact of the brain, contact of the b
                                                                                                                                                       Kit for diagnozing and treating T-cell associated diseases e.g. autoimmune, degenerative nervous system and infectious disease, comprises nucleic acid primers specifically priming and allowing amplification of a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              breast. The present sequence represents a Vbeta gene repeat sequence.
                                                                                                                                                                                                                                                                                                                                                                 Disclosure; SEQ ID NO 596; 164pp; English.
                                                                           WPI; 2004-059052/06
                                                                                                                                                                                                                                                                                     Vbeta gene
```

Score 15.8; DB 1; Length 20; Pred. No. 2e+02; 0; Mismatches 2; Indels Ze+02; ~hes 2; Indels Sequence 20 BP; 8 A; 0 C; 0 G; 12 T; 0 U; 0 Other; 3300 ATATTTTTTTTTATA 3318 Arararrrarrarara 20 0.4%; Local Similarity 89.5 hes 17; Conservative Query Match Matches ò 셤

o;

Gaps

°

ADH67733 standard; DNA; 20 25-MAR-2004 (first entry) ADH67733; RESULT 178 ADH67733 

BP

Human glucocorticoid receptor-specific antisense oligonucleotide #4567.

antisense oligonucleotide; glucocorticoid receptor; infection; inflammation; tumour formation; diabetes; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2'-methoxyethyl; 2'-MOE.

Homo sapiens

WO2003099215-A2.

04-DEC-2003.

20-MAY-2003; 2003WO-US016084.

20-MAY-2002; 2002US-0381857P.

(PHAA ) PHARMACIA CORP

to nucleic acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence

The invention comprises an antisense oligonucleotides that are targeted

Nalseth AE Crosby SD,

WPI; 2004-035034/03.

```
ö
                                                                                                             The invention comprises an antisense oligonucleotides that are targeted to nucleic acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-NOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
                             mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            antisense oligonucleotide; glucocorticoid receptor; infection; inflammation; tumour formation; diabetes; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2'-methoxyethyl; 2'-MOS.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Human glucocorticoid receptor-specific antisense oligonucleotide #2769
                                                                                                                                                                                                                                                                                                                                                       Gaps
               New antisense compound targeted to a nucleic acid molecule encoding
                                                                                                                                                                                                                                                                                                                                                     ;
0
                                                                                                                                                                                                                                                                                                                0.4%; Score 15.8; DB 1; Length 20; 89.5%; Pred. No. 2e+02; ive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                      Sequence 20 BP; 12 A; 1 C; 0 G; 7 T; 0 U; 0 Other;
                                                                                 ID NO 4567; 985pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Claim 4; SEQ ID NO 2769; 985pp; English
                                                                                                                                                                                                                                                                                                                                                                                       3298 TGATATTTTTTTA 3316
                                                                                                                                                                                                                                                                                                                                                                                                                        N
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADH65935 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    20-MAY-2002; 2002US-0381857P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20-MAY-2003; 2003WO-US016084.
                                                                                                                                                                                                                                                                                                                                                                                                               20 TAATATTTTTATATTTA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (first entry)
                                                                                                                                                                                                                                                                                                                                                       17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Crosby SD, Nalseth AE,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (PHAA ) PHARMACIA CORP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2004-035034/03.
                                                                                                                                                                                                                                                                                                                                         Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WO2003099215-A2.
                                                                                  Claim 4; SEQ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              25-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     04-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADH65935;
                                                                                                                                                                                                                                                                                                                       Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 179
                                                                                                                                                                                                                                                                                                                                                         Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADH65935
à
                                                                                                                                                                                                                                                                                                                                                                                                                        d
```

10001863-3.81.rng

(first entry)

```
The invention relates to novel compounds of 8-80 nucleobases in length targeted to, and which specifically hybridises with, a nucleic acid molecule encoding 3-hydroxy-3-methylglutaryl-Consyme A (HMG-COA) reductase, and inhibits the expression of HMG-COA reductase. The novel compounds have cardiant, antiarteriosclerotic, and antilipaemic activities. The compound can be used to treat disorders by antisense gene therapy. The compounds, compositions and methods are useful for treating a disease or condition associated with HMG-COA reductase, such as a cardiovascular disorder e.g. atherosclerosis, or a disease or condition
                                                                                                                                         HMG-CoA reductase; 3-hydroxy-3-methylglutaryl-Coenzyme A;
HMG-CoA reductase; cardiant; antiarteriosclerotic; antilipaemic;
antisense gene therapy; cardiovascular disorder; cholesterol metabolism;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      New compounds, particularly antisense oligonucleotides targeted to
                                                                                                        Human HMG-CoA reductase antisense oligonucleotide, SEQ ID No 248
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        nucleic acid encoding HMG-COA reductase, useful for treating atherosclerosis, or a disease involving cholesterol metabolism
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Example 16; SEQ ID NO 248; 110pp; English.
   ADI79725 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                         32-JUL-2002; 2002US-00190366.
                                                                                                                                                                                                                                                                                                                                                                            02-JUL-2002; 2002US-00190366.
                                                                                                                                                                                                                                                                                                                                                                                                               (ISIS-) ISIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                  Dean NM, Freier SM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 VPI; 2004-081743/08
                                                                                                                                                                                                                                                                   US2004006031-A1.
                                                                                                                                                                                                                                    Homo sapiens,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           angiogenesis.
                                                                       22-APR-2004
                                                                                                                                                                                                                                                                                                      08-JAN-2004.
                                                                                                                                                                                                 human; 88.
   a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ò
                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           The invention comprises an antisense oligonucleotides that are targeted antisense acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothicate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                           antisense oligonucleotide; glucocorticoid receptor; infection; inflammation; tumour formation; diabetees; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2 - methoxyethyl; 2 '-MOE.
                                                                                                                                                                                                                                                                                                                                                                          Human glucocorticoid receptor-specific antisense oligonucleotide #4537.
                                                                                                          Gaps
                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15.8; DB 1; Length 20; 9.5%; Pred. No. 2e+02; ve 0; Mismatches 2; Indels
                                                                       DB 1; Length 20;
                                                                                                       2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 20 BP; 13 A; 0 C; 1 G; 6 T; 0 U; 0 Other;
                                 Sequence 20 BP; 5 A; 3 C; 5 G; 7 T; 0 U; 0 Other;
                                                                                    2e+02;
                                                                  Score 15.8; DB; Pred. No. 2e+02 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Claim 4; SEQ ID NO 4537; 985pp; English.
                                                                                                                                         1325 CAAAGGTTGCTGTTCTCAA 1343
                                                                                                                                                                         1 CAAATGTTGCTGTTCTGAA 19
                                                                                                                                                                                                                                                                 ADH67703 standard; DNA; 20 BP
                                                                       0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20-MAY-2003; 2003WO-US016084.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20-MAY-2002; 2002US-0381857P.
                                                                                                                                                                                                                                                                                                                                       25-MAR-2004 (first entry)
                                                              Query Match
Best Local Similarity 89.5
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Crosby SD, Nalseth AE;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (PHAA ) PHARMACIA CORP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 2004-035034/03.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match
Best Local Similarity
Matches 17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WO2003099215-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         04-DEC-2003
                                                                                                                                                                                                                                                                                                    ADH67703;
                                                                                                                                                                                                                                RESULT 180
                                                                                                                                                                                                                                                  ADH67703,
                                                                                                                                                                                                                                                                 ઠ
                                                                                                                                                                         셤
```

Ø

₹

Dobie

```
ö
cardiovascular disorder e.g. atherosclerosis, or a disease or condition involving cholesterol metabolism. They are also useful in research and diagnostics for modulating the expression of HWG-CoA reductase. This polynucleotide sequence represents an antisense oligonucleotide of the
                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            HMG-CoA reductase; 3-hydroxy-3-methylglutaryl-Coenzyme A;
HMG-CoA reductase; cardiant; antiarteriosclerotic; antilipaemic;
                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human HMG-CoA reductase antisense oligonucleotide, SEQ ID No
                                                                                                                                                Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                              Sequence 20 BP; 4 A; 4 C; 9 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                           73 CICTGATCCCAGCCATGGC 91
                                                                                                                                                                                                                                                                crarecreceaecearese 1
                                                                                                                                                                                                                                                                                                                                                            ВP
                                                                                                                                                   0.4%;
                                                                                                                                                                                                                                                                                                                                                            ADI79528 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                      (first entry)
                                                                                                                                                                                      Conservative
                                                                                                                                                                   Similarity
17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                      22-APR-2004
                                                                                                                                                                                                                                                                19
                                                                                                                                                                                                                                                                                                                                                                                                 ADI 79528;
                                                                                                                                                   Query Match
                                                                                                                                                                         Local
                                                                                                                                                                                                                                                                                                                     RESULT 182
ADI79528
                                                                                                                                                                                        Matches
```

Gaps

ö

3297 CTGATATATTTTTATTTT 3315

8 셤

89.58;

Conservative

19

ADI 79725/c RESULT 181

```
25-JUL-2003; 2003WO-US023509.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (EPIG-) EPIGENESIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADJ597:
    g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The invention relates to novel compounds of 8-80 nucleobases in length targeted to, and which specifically hybridises with, a nucleic acid molecule encoding 3-hydroxy-3-methylglutaryl-Coneryme A (HMG-COA) reductase, and inhibits the expression of HMG-COA reductase. The novel compounds the used to treat disorders by antisense gene therapy. The compound can be used to treat disorders by antisense gene therapy. The compound can be used to treat disorders by antisense gene therapy. The compounds compositions and methods are useful for treating a disease or condition associated with HMG-COA reductase, such as a cardiovascular disorder e.g. atherosclerosis, or a disease or condition involving cholesterol metabolism. They are also useful in research and diagnostics for modulating the expression of HMG-COA reductase. This polymucleotide sequence represents an antisense oligonucleotide of the
antisense gene therapy; cardiovascular disorder; cholesterol metabolism;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 interleukin; IL-4 receptor; IL-5 receptor; lung disease; airway inflammation; allergy; asthma; impeded respiration; cystic fibrosis; acute respiratory distress syndrome; pulmonary hypertension; lung inflammation; bronchitis; oligonucleotide;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                  ಗ
                                                                                                                                                                                                                                                                                                                                                                                                                           New compounds, particularly antisense oligonucleotides targeted to
nucleic acid encoding HMG-CoA reductase, useful for treating
atherosclerosis, or a disease involving cholesterol metabolism or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 3 A; 9 C; 4 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Example 15; SEQ ID NO 51; 110pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Oligonucleotide associated to PDE4A #86
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          CTCTGATCCCAGCCATGGC 91
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      crariscricicascicarissic 20
                                                                                                                                                                                                    02-JUL-2002; 2002US-00190366.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      0.4%;
                                                                                                                                                                                                                                                02-JUL-2002; 2002US-00190366
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ADJ60803 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             06-MAY-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Local Similarity 89.5 tes 17; Conservative
                                                                                                                                                                                                                                                                                             (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                       Freier SM,
                                                                                                                                                                                                                                                                                                                                                                                   WPI; 2004-081743/08
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WO2004011613-A2
                                                                                                             US2004006031-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 angiogenesis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Homo sapiens.
                                                                      Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            05-FEB-2004.
                                                                                                                                                          08-JAN-2004
                         human; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          73
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ADJ60803;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
                                                                                                                                                                                                                                                                                                                                       Dean NM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ADJEONALT 183
ADJEONAGO ON ACCOUNT 183
ACCOUNT OF ADJEONAL ACCOUNT ON ACCOUNT
8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  g
```

```
The present invention relates to an oligomucleotide anti-sense to e.g., initiation codon, coding region with 2-10 nucleotides of 5'-end and 3'-end of nucleic acid target comprising gene(s) chosen from e.g. interleukin (IL)-4 receptor, IL-5 receptor or salts of the oligomucleotide and optionally surfactant operatively linked to the oligomucleotide. The method is useful for preventing or treating a cligomucleotide the method is useful for preventing or treating a ceptratory or lung disease, which involves administering to the airways of a subject an effective amount of an inhibitor. The oligomucleotide is useful for production of a medicament for the prevention and/or treatment useful for production of a medicament for the prevention and/or treatment cof a respiratory or lung disease. The respiratory or lung disease is chosen from airway inflammation, allergy(ies), asthma, impeded repiration, cystic fibrosis (CF), chronic obstructive pulmonary diseases (CRDS), allergic rhinitis (AR), acute respiratory distress syndrome (ARDS), pullmonary hypertension, lung inflammation, bronchitis, airway obstruction. The present sequence represents an oligomucleotide of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                     Novel single or multiple target oligonucleotide anti-sense to e.g. initiation codons and introns of respiratory disease-relevant genes e.g., CCR1, RANTES, MCP4, useful for prophylaxis or treating respiratory
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             interleukin; IL-4 receptor; IL-5 receptor; lung disease; airway inflammation; allergy; asthma; impeded respiration; cystic fibrosis; acute respiratory distress syndrome; pulmonary hypertension; lung inflammation; bronchitis; oligonucleotide;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.4%; Score 15.8; DB 1; Length 20;
89.5%; Pred. No. 2e+02;
ive 0; Mismatches 2; Indels
                                                                                                                                      Miller S;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 3 A; 6 C; 8 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Oligonucleotide associated to Eotaxin U46572 #5.
                                                                                                                                  Aguilar D,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Claim 2; SEQ ID NO 1659; 85pp; English.
                                                                                                                                         Sandrasagra A,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    2508 CAGGTGGAGCTGTACCGCC 2526
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ~
                                                                                                                                                                   Cong H;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           29-JUL-2002; 2002US-0399076P.
29-JUL-2002; 2002US-0399076P.
                                                                    (EPIG-) EPIGENESIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ccecreeaecreraccecc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ADJ59719 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   17; Conservative
                                                                                                 Tang L, c...
Lu H,
                                                                                                                                                                                                                                                                                                                                                                                                                         disease e.g., asthma
                                                                                                                                                                                                                                           WPI; 2004-203534/19.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO2004011613-A2
                                                                                                                                                                          Shahabuddin S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          25-JUL-2003;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              06-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     05-FEB-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               nvention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ADJ59719;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Best Local
                                                                                                                                         Nyce JW,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               RESULT 184
```

```
Disclosure; SEQ ID NO 34; 56pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 3766 TGGCTGGGATCCCTCCCCT 3784
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20 réderédearcérereir 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    20-AUG-2003; 2003WO-GB003637
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            24-AUG-2002; 2002GB-00019890.
26-AUG-2002; 2002US-0405717P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               89.24;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                COLLEGE LONDON
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ADL23352 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Conservative
                            WPI; 2004-132608/13
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2004-226867/21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Local Similarity
tes 17; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WO2004018711-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (UNICO ) UNIV
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Ming-Qing D;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      04-MAR-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADL23352;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 186
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADL23352/
    ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           The present invention relates to an oligonucleotide anti-sense to e.g., initiation codon, coding region with 2-10 nucleotides of 5'-end and 3'-end of nucleic acid target comprising gene(s) chosen from e.g. and of nucleic acid target comprising gene(s) chosen from e.g. interleukin (IL)-4 receptor, IL-5 receptor or salts of the cligonucleotide and optionally surfactant operatively linked to the cligonucleotide. The method is useful for preventing or treating a respiratory or lung disease, which involves administering to the airways of subject an effective amount of an inhibitor. The oligonucleotide is useful for production of a medicament for the prevention and/or treatment of a respiratory or lung disease. The respiratory or lung disease is chosen from airway inflammation, allergy(iss), asthma, impeded consen from airway inflammation, allergy(iss), asthma, impeded consen from airway inflammation, allergy (iss), asthma, impeded consenting chinitis (AR), acute respiratory distress syndrome (ARDS), pulmonary hypertension, lung inflammation, bronchitis, airway obstruction. The present sequence represents an oligonucleotide of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                              e.g.,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                Novel single or multiple target oligonucleotide anti-sense to e.g. initiation codons and introns of respiratory disease-relevant genes CCR1, RANTES, MCP4, useful for prophylaxis or treating respiratory
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     0.4%; Score 15.8; DB 1; Length 20;
89.5%; Pred. No. 2e+02;
ative 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              oligonucleotide, human, B7 protein, B7-1; asthma, atic, antiinflammatory, 88.
                       Nyce JW, Tang L, Sandrasagra A, Aguilar D, Miller S;
Shahabuddin S, Lu H, Cong H;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Airway hyperresponsiveness; pulmonary inflammation;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence 20 BP; 3 A; 7 C; 4 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Human B7-1 DNA antisense oligonucleotide #25.
                                                                                                                                                                                                                                              Claim 2; SEQ ID NO 575; 85pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Bennett CF, Vickers TA, Karras JG;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                2707 GAGGCATTTCTTGCCCAGC 2725
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 1 GAGGCATTTCTTGTCCACC 19
                                            Cong H;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADJ54227 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             31-DEC-1996; 96US-00777266.
04-UUN-1999; 99US-00326186.
25-WAY-2000; 2000WO-US014471.
09-WAY-2001; 2001US-00851871.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   23-MAY-2003; 2003US-00444206
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 06-MAY-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Local Similarity 89.5
nes 17; Conservative
                                                                                                                                                                                                      disease e.g., asthma.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             BENNETT C F.
VICKERS T A.
                                                                                        WPI; 2004-203534/19.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (VICK/) VICKERS T A (KARR/) KARRAS J G.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            US2004023917-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 antisense oligo
antiasthmatic;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      05-FEB-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ADJ54227;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (BENN/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADJ54227,
PX A P B X X P X P X P X S X X W W X X D X A X D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D X Y D
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          g
```

```
ö
                                                                                                                                                                                                                                                                               The invention relates to a method for treating airway hyperresponsiveness or pulmonary inflammation in an individual comprising administering an antisense compound targeted to a nucleic acid molecule encoding a human B7 protein. The invention also relates to a method of inhibiting
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ss; primer; diagnosis; cervical intraepithelial neoplasia; CIN; allelic deletion; FHIT; fragile histidine triad gene; PR; progesterone receptor; DLBC1; deleted in lung and oesophageal cancer 1; TRIM29; tripartite motif-containing 29; microsatellite; D3S1300; D3S1260; D1SS35; D1ISS28.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Diagnosing cervical intraepithelial neoplasia comprising detecting an allelic deletion in genes selected from FHIT, PR, DLEC1- or TRIM 29 by comparing the FHIT, PR, DLEC1 and/or TRIM 29 polynucleotides or proteins present in the samples.
Treating airway hyperresponsiveness or pulmonary inflammation comprises administering an antisense compound targeted to a nucleic acid molecule encoding a human B7 protein to the individual.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               expression of a nucleic acid molecule encoding B7-1 or B7-2. The antisense compound is an antisense oligonucleotide which has a modified sugar moiety and nucleobase. The human B7 protein is human B7-1 or B7-2 protein or both. The compound is useful for treating alrway
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   hyperresponsiveness or pulmonary inflammation, which is associated with asthma, by inhibiting expression of human B7 protein. This sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%; Score 15.8; DB 1; Length 20; 39.5%; Pred. No. 2e+02; ve 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        represents an antisense oligonucleotide of the invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 20 BP; 7 A; 4 C; 8 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                               Example 1; SEQ ID NO 47; 182pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Primer #2 for identification of HPV 45.
```

```
This sequence represents a primer which was used in the method of the invention for diagnosing susceptibility to persistence or progression of cervical intraspithelial neoplasia (CIN) in an individual suffering from the disease. The method comprises detecting an allelic deletion in one or compares selected from FHIT (fragile histidine triad gene), PR (progrestorone receptor), DLECI (deleted in lung and ossophageal cancer 1) or TRIM29 (tripartite motif-containing 29) by comparing the FHIT, PR, DLECI and/or TRIM29 polynucleotides or proteins present in the samples contributed from non-dyskaryotic and dyskaryotic samples, respectively. The method is carried out using a kit comprising a panel of two or more pairs of primers, where each pair of primers is suitable for amplifying a microsatellite DNA marker selected from DS31260, D351260, D1835 or D118528, or a panel of two or more specific binding agents, where each contributed distinguishing between the normal and allelic deletion forms of a polynucleotide or protein selected from FHIT, PR, TRIM29 or DLECI. The method is useful for diagnosing susceptibility to persistence or progression of cervical intraeplihalial neoplasia in an and individual suffering from the disease.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Antisense therapy; human; Fas; Fas ligand; FasL; Apo-1L; CD95L; Fas associated protein 1; Fap-1; signal transduction; autoimmune disease; inflammatory disease; cancer; immunosuppressive; antiinflammatory; cytostatic; phosphorothioate; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        New antisense oligonuclectides of 20-50 nucleobases, useful for treating autoimmune or inflammatory diseases, and cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         /mote= "This oligonuclectide has a phosphorothioate backbone and 2-methyoxyethyl (2' MOS) wings at the 5' and 3' ends, which are 5 nuclectides in length at each end. All cytidine residues are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 1 A; 4 C; 6 G; 9 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Human Fas cDNA, antisense oligonucleotide #72.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1995 ACACCTTCAGATAAGCAGG 2013
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Wyatt J;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20 ACACCTCCAGAAAAGCAGG 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 18-SEP-2000; 2000US-00665615.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      99US-00290640.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Match 0.4%;
Local Similarity 89.5%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADL27792 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ø
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20-MAY-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Dean NM, Marcusson EG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               _bom/
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WPI; 2004-050524/05.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  US6653133-B1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          12-APR-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                25-NOV-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ADL27792;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 18
ADL27792/
XX
AC ADL2
XX
XX
ADL2 20-M
DT 20-M
                     ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  d
```

```
The present invention relates to antisense compounds targeted to nucleic acids encoding human Fas (also known as Apo-1 or CD95), Fas ligand (FasL, also Apo-1L and CD951), and Fas associated protein 1 (Fap-1). The antisense compound comprises an antisense oligonucleotide that specifically hybridises with one of the said nucleic acids and inhibits Fas, FasL or Fap-1 mediated signal transduction. The antisense oligonucleotide is a chimeric oligonucleotide. The antisense oligonucleotide comprises at least one modified internucleoside linkage, preferably a phosphorothioate linkage. It also comprises at least one modified anciented antisense oligonucleotide further comprises at least one modified nucleobase, preferably a 5-methylcytosine. The antisense oligonucleotide further comprises at least one modified nucleobase, preferably a 5-methylcytosine. The antisense oligonucleotide further comprises at least one modified matery diseases, and cancers associated with overexpression of constitutive activation of Fas, FasL, or Fap-1. The present sequence
                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         immunosuppressive; antiinflammatory; hepatotropic; virucide; cytostatic; antisense technology; Fas; Fas ligand; Fap-1; Fas associated disorder; pap-1 associated disorder; ischaemia reperfusion injury; apoptosis; allograft; autteimmune disease; inflammatory disease; hepatitis; cancer; lymphoma; human; antisense oligonucleotide; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       /mod_base= OTHER
/note= "OTHER= Phosphorothioate backbone. All cytidines
are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /*tag= a
/mod_base= OTHER
/note= "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /mod_base= OTHER
/note= "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                  .;
0
                                                                                                                                                                                                                                                                                                                                                                              Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                             Sequence 20 BP; 9 A; 1 C; 0 G; 10 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Human Fas antisense oligonucleotide seqid 153.
Example 18; SEQ ID NO 153; 76pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                    861 TTAAGAAATAATTTGATA 879
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20 TTAAGAAATAATATTATA 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      12-APR-1999; 99US-00290640.
18-SEP-2000; 2000US-00665615.
09-MAR-2001; 2001US-00802669.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    14-JUL-2003; 2003US-00619220.
                                                                                                                                                                                                                                                                                                                                                                                0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADM53564 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Д
                                                                                                                                                                                                                                                                                                                                                                            Query Match
Best Local Similarity 89.5'
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            15. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1. .20
/*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /*tag=
                                                                                                                                                                                                                                                                                                                present invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   US2004033979-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              03-JUN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     19-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADM53564;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RESULT 188
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADM53564/
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      g
                                                                                                                                                                                                                                                                                                                                                                                                       .
0
```

```
The invention describes an antisense compound 8-30 or 8-50 nucleobases in length targeted to the 5'-untranslated region, translational start site, translational termination region or 3'-untranslated region of a nucleic acid molecule encoding Fas, Fas ligand or Fap-1. Also described are: a pharmaceutical composition comprising the anti-sense compound and a pharmaceutical carrier or diluent, a method of inhibiting the expression of Fas or Fap-1 in cells or tissues: treating an animal having a disease or condition associated with Fas or Fap-1; anda preventing allograft recipient. The antisense compound and pharmaceutical composition is useful in diagnosing, treating or preventing autoimmune or inflammatory disease. G. hepatitis, cancer, e.g. cancer of the colon, liver, lung or a lymphoma, apoptosis, allograft rejection, e.g. cardiac, real, hepatic or skin allograft rejection, injury. This sequence
                                                                                                                                                                                                 New antisense compound targeted to nucleic acid molecule encoding Fas or Fap-1, useful in diagnosing, treating or preventing autoimmune or inflammatory disease, cancer, apoptosis, allograft rejection or ischemia reperfusion injury.
                                                                                                                                                                                                                                                                                                          Example 18; SEQ ID NO 153; 83pp; English
                                                                                                                        Dean NM, Marcusson EG,
                   DEAN N M.
MARCUSSON E G.
WYATT J.
ZHANG H.
                                                                                                                                                              WPI; 2004-180091/17.
                 (DEAN/) 1 (MARC/) 1 (WYAT/) 1
                                                                               ZHAN/)
```

Gaps ö 0.4%; Score 15.8; DB 1; Length 20; 89.5%; Pred. No. 2e+02; attive 0; Mismatches 2; Indels Sequence 20 BP; 9 A; 1 C; 0 G; 10 T; 0 U; 0 Other; represents a human Fas antisense oligonucleotide Query Match
Best Local Similarity 89.5'
Matches 17; Conservative

ö

861 TTAAGAAATAATTTTGATA 879 rraagaaraararrara 2 20 RESULT 189 ઠે 유

ADO45209 standard; DNA; 20 Human oligonucleotide #575. 15-JUL-2004 (first entry) ADO45209; AD045209 

Human; ss; interleukin-4 receptor; IL-4; interleukin-5 receptor; IL-5; CCR1; CCR3; Ectaxin-1; RANTES; MCP4; CD23; ICAM; UCAM; tryptase a; tryptase b; PDE4 A; PDE4 B; PDE4 C; PDE4 D; respiratory disease; lung disease; hyper-responsiveness; adenosine; adenosine A receptor; asthma; lung allergy; inflammation; inflammatory disease; alway inflammation; allergy; impeded respiration; cystic fibrosis; CF; chronic obstructive pulmonary disease; COPD; allergic rhinitis; acute respiratory distress syndrome; pulmonary hypertension; lung inflammation; bronchitis; airway obstruction; bronchoconstriction.

Homo sapiens

US2004049022-A1

11-MAR-2004.

25-JUL-2003; 2003US-00627930

23-APR-2002; 2002WO-US013135

23-APR-2002; 2002WO-US013143.

NYCE J W. SANDRASAGRA (NYCE/) (SAND/) (TANG/) (TANG/) (AGUI/) (MILL/) (SHAH/) (LUHH/) (CONG/) (CONG/)

TANG L.
AGUILAR D.
MILLER S.
SHAHABUDDIN S.

Zhang H;

Wyatt J,

CONG H.

Sandrasagra A, Tang L, Aguilar D, Miller S; In S, Lu H, Cong H; Shahabuddin S, Lu H, Nyce JW,

WPI; 2004-293804/27.

Novel single or multiple target oligonucleotide anti-sense to e.g. initiation codon, intron of respiratory disease-relevant gene e.g. CCR1, RANTES, MCP4, useful for prophylaxis or treating respiratory disease e.g. asthma.

Claim 2; SEQ ID NO 575; 174pp; English

The invention relates to oligonuclectides anti-sense to an initiation codon, coding region, 5' or 3' intron-exon junction, intron or region to codon, coding region, 5' or 3' intron-exon junction, intron or region codon, coding region, 5' or 3' intron-exon junction, intron or region codon, coding interleukin (11.0-4 receptor, interleukin (11.0-5) receptor, CCR1, CCR3, Eotaxin-1, RANTES, MCP4, CD23, ICAM, VCAM, tryptase a, tryptase b, PDE4 A, PDE4 B, PDE4 C or PDE4 D. The invention also relates to a method of screening a candidate compound that binds to one or more nucleic acid target(s) or expressed product(s), for the prevention and/or treatment of a respiratory or lung disease. The cligonucleotides are useful for reducing or inhibiting expression of a cligonucleotides are useful for reducing or inhibiting expression of a creapiratory or lung disease. The creapiratory or lung disease are useful for preventing or treating a respiratory or lung disease. The creapiratory or lung disease is associated with hyper-responsiveness to cadhor increased levels of, adenosine and/or lung allergie associated with creapiratory or lung disease conformation or an inflammation, allergy, asthma, impeded respiration, cystic fibrosis (CF), chronic obstructive pulmonary disease (CC is chosen from airway inflammation, allergy, asthma, impeded respiration, cystic fibrosis (CF), chronic obstructive pulmonary disease (CDPD), contractive and contractive pulmonary disease (CF) thouse of the proper and contractive pulmonary disease (CPPD) thouse the contractive pulmonary disease (CPPD) the contractive pulmonary disease (CPPD), chronic obstructive pulmonary disease (CPPD) the contractive descriptor or the contractive descriptor or the contractive descriptor o pronchoconstriction. This sequence represents an oligonucleotide of 

Sequence 20 BP; 3 A; 7 C; 4 G; 6 T; 0 U; 0 Other;

Gaps ö 0.4%; Score 15.8; DB 1; Length 20; 89.5%; Pred. No. 2e+02; ative 0; Mismatches 2; Indel8 Matches 17; Conservative Local Similarity Query Match

ö

ð

ADO46292 standard; DNA; 20 BP. 15-JUL-2004 (first entry) ADO46292; ADO46292/ 

RESULT 190

Human; 88; interleukin-4 receptor; IL-4; interleukin-5 receptor; IL-5; CCR1; CCR3; BCCAXin-1, RAPTES; MCP4; CD3; ICR4), VCR4), tryptase a; tryptase CCR3; DCR4 PDE4 B; PDE4 C; PDE4 D; respiratory disease; tryptase; hyper-responsiveness; adenosine; adenosine A receptor; Human oligonucleotide #1658.

ADAMIS; metagirdin; MDCIS; a disintegrin and metalloproteinase domain 15; diagnosis; inflammation; therapy; human; ss.

Human ADAM15 target oligonucleotide #26.

(first entry)

12-AUG-2004

ADO51802;

BP.

ADO51802 standard; DNA; 20

RESULT 191

ADO51802

```
asthma, lung allergy; inflammation; inflammatory disease; airway inflammation; allergy; impeded respiration; cystic fibrosis; CF; chronic obstructive pulmonary disease; COPD; allergic rhinitis; acute respiratory distress syndrome; pulmonary hypertension; lung inflammation; bronchitis; airway obstruction; bronchoconstriction.
                                                                                                                                                                                                                                                                                                            Novel single or multiple target oligonucleotide anti-sense to e.g. initiation codon, intron of respiratory disease-relevant gene e.g. CC RANTES, MCP4, useful for prophylaxis or treating respiratory disease
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence 20 BP; 3 A; 6 C; 8 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                           Claim 2; SEQ ID NO 1659; 174pp; English.
                                                                                                                                                                                                                                                            Nyce JW, Sandrasagra A, Tang L,
Shahabuddin S, Lu H, Cong H;
                                                                                                                     25-JUL-2003; 2003US-00627930.
                                                                                                                                         23-APR-2002; 2002WO-US013135.
23-APR-2002; 2002WO-US013143.
                                                                                                                                                                      NYCE J W.
SANDRASAGRA A.
                                                                                                                                                                                                            MILLER S.
SHAHABUDDIN S.
                                                                                                                                                                                                                                                                                         WPI; 2004-293804/27
                                                                                                                                                                                        TANG L.
AGUILAR D.
                                                                              US2004049022-A1
                                                                                                                                                                                                                                         CONG H.
                                                            Homo sapiens.
                                                                                                  11-MAR-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
                                                                                                                                                                     (NYCE/)
(SAND/)
                                                                                                                                                                                                                                         (CONG/)
                                                                                                                                                                                                             (MILL)
                                                                                                                                                                                                                     SHAH/)
                                                                                                                                                                                                                                                                                                                                            asthma.
                                                                                                                                                                                                                               LUHH/
                                                                                                                                                                                        (TANG/)
```

The invention relates to oligonucleotides anti-sense to an initiation codon, coding region, 5' or 3' intron-exon junction, intron or region with 2-10 mucleotides of the 5'-end or 3'-end of a mucleic acid target chosen from a gene encoding interleukin (IL)-4 receptor, interleukin (IL)-5 receptor, CCR1, CCR3, Ectaxin-1, RANTES, MCP4, CD23, ICAM, VCAM, tryptase a, tryptase b, PDB4 A, PDB4 B, PDB4 C or PDB4 D. The invention also relates to a method of screening a candidate compound that binds to one or more nucleic acid target(s) or expressed product(s), for the prevention and/or treatment of a respiratory or lung disease. The cligonucleotides are useful for reducing or inhibiting expression of a gene or mRNA encoding interleukin-5 receptor, CCR1, CCR3, Ectaxin-1, RANTES, MCP4, CD23, ICAM, VCAM, tryptase a, tryptase b, PDB4 B, PDB4 C, or PDB4 D. The oligonucleotides are useful for preventing or treating a respiratory or lung disease. The respiratory or lung disease is associated with hyper-responsiveness to and/or increased levels of, adenosine and/or levels of adenosine A receptor(8), and/or asthma and/or lung allergies associated with inflammation or an inflammatory disease. The respiratory or lung disease is chosen from airway inflammation, allergy, asthma, impeded respiration, cystic fibrosis (CF), chronic obstructive pulmonary disease (COPD), allergic rhinitis, acute respiratory distress syndrome, pulmonary hypertension, lung inflammation, bronchitis, airway obstruction or bronchoconstriction. This sequence represents an oligonucleotide of the

ö

to ADAMIS (otherwise known as metagirdin, MDC15, and a disintegrin and metalloproteinase domain 15) and which modulate the expression of ADAMIS. The invention is useful for diagnosing and treating diseases associated with expression of ADAMIS such as inflammation. The present sequence is human ADAMIS target oligonucleotide. This sequence is used in the exemplification of the invention.

Sequence 20 BP; 6 A; 5 C; 6 G; 3 T; 0 U; 0 Other;

The present invention is directed to antisense oligonucleotides targeted

Example 15; SEQ ID NO 73; 38pp; English.

e.g

CCR1

New compound targeted to a nucleic acid molecule encoding ADAM15 and inhibits the expression of ADAM15, useful for modulating the expression of ADAM15 or for diagnosing or treating, e.g. inflammation.

Dobie KW;

WPI; 2004-399722/37.

PHARM INC Dean NM,

SISI (-SISI)

Bennett CF,

Miller S;

Aguilar D,

21-NOV-2002; 2002US-00302028. 21-NOV-2002; 2002US-00302028

US2004102392-A1.

27-MAY-2004.

Homo sapiens.

```
ADAWIS; metagirdin; MDCIS; a disintegrin and metalloproteinase domain 15; diagnosis; inflammation; therapy; human; antisense; phosphorothioate backbone; ss.
                         Gaps
                         ;
0
0.4%; Score 15.8; DB 1; Length 20; 89.5%; Pred. No. 2e+02; ative 0; Mismatches 2; Indels
                                                                                                                                                                                                     Human ADAM15 antisense oligonucleotide ISIS #173654.
                                                                                                                                                                                                                                                                                                                Location/Qualifiers
                                                 2601 AAAGCCCTGCTGGATGGTA 2619
                                                                    2 AAAGCCCTCCTGGATGGAA 20
                                                                                                                                   ADO51767 standard; DNA; 20 BP.
                                                                                                                                                                                (first entry)
                           17; Conservative
 Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                            Homo sapiens.
                                                                                                                                                                                  12-AUG-2004
                                                                                                                                                                                                                                                                                         Synthetic
                                                                                                                                                          ADO51767;
                                                                                                            RESULT 192
                         Matches
                                                                                                                      ò
                                                                       셤
                                                                                                                                                                                                                                                        ;
0
                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                        ö
```

0.4%; Score 15.8; DB 1; Length 20;

Pred. No. 2e+02; 0; Mismatches

89.58;

Conservative

Local Similarity nes 17; Conserv

Best Loca Matches

2508 CAGGTGGAGCTGTACCGCC 2526

ઠે 셤

N

ccecreascreracecc

20

Indels

5

10001863-3.sl.rng

'mod base= OTHER

```
20 cagccacacccrcrccrcrcr
                                                                                                                                  27-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Matches
à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    g
                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                /mod_base= OTHER
/note= "Phosphorothioate backbone where all cytidines are
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         cytostatic; antisense therapy; cytokine-inducible kinase; cytokine-inducible kinase inhibitor; antisense technology; cytokine-inducible kinase expression; hyperproliferative disorder; human;
                                                                                                                                                                                                                                                                                                                          The present invention is directed to antisense oligonucleotides targeted to ADAM15 (otherwise known as metagirdin, MDC15, and a disintegrin and metalloproteinase domain 15) and which modulate the expression of ADAM15.
                                                                                                                                                                                                                                                                   New compound targeted to a nucleic acid molecule encoding ADAMIS and inhibits the expression of ADAMIS, useful for modulating the expression of ADAMIS or for diagnosing or treating, e.g. inflammation.
                                                                                                                                                                                                                                                                                                                                                        The invention is useful for diagnosing and treating diseases associated with expression of ADAMIS such as inflammation. The present sequence is human ADAMIS antisense oligonucleotide. This sequence is used in the
                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                               /mod_base= OTHER
/note= "2' -methoxyethyl (2' -MOE) nucleotides"
16. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                              ô
                                                                                                               'note= "2' -methoxyethyl (2' -MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Human cytokine-inducible kinase antisense oligonucleotide #30.
                                                                                                                                                                                                                                                                                                                                                                                                                        0.4%; Score 15.8; DB 1; Length 20; 89.5%; Pred. No. 2e+02; Live 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 3 A; 6 C; 5 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                         Example 15; SEQ ID NO 38; 38pp; English.
                                     5-methyl cytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                      exemplification of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                               2601 AAAGCCCTGCTGGATGGTA 2619
                                                                                             /*tag= c
/mod base= OTHER
                                                                                                                                                                                                                               Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAAGCCCTCCTGGATGGAA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ВЪ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        antisense oligonucleotide; ss
                                                                                                                                                                      21-NOV-2002; 2002US-00302028.
                                                                                                                                                                                         21-NOV-2002; 2002US-00302028
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 193
ADN30059/c
ID ADN30059 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Ω
       *tag= b
                                                        ๙
                                                                                                                                                                                                                                                                                                                                                                                                                                              Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1. .20
/*tag=
                                               . 5
*tag≃
                                                                                                                                                                                                            (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                               Dean NM,
                                                                                                                                                                                                                                                 WPI; 2004-399722/37.
                                                                                                                                                                                                                                                                                                                                                                                                                                   Local Similarity
                                                                                                                                  US2004102392-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      modified_base
modified base
                                             modified_base
                                                                                  modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     12-AUG-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Homo sapiens
                                                                                                                                                                                                                              Bennett CF,
                                                                                                                                                                                                                                                                                                                                                                                                                                            17;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  19
                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADN30059
                                                                                                                                                                                                                                                                                                                                                                                                                                     Best Loca
Matches
셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ð
```

```
The invention describes a compound 8-80 nucleobases in length targeted to a nucleic acid molecule encoding Cytokine-inducible kinase. The compound specifically hybridises with the nucleic acid molecule encoding Cytokine-inducible kinase (which comprises a sequence of 2169 bp fully defined in the described are: a method of inhibiting the expression of thinase. Also described are: a method of inhibiting the expression of Cytokine-inducible kinase in cells or tissues, comprising contacting the cells or tissues with the new compound so that the expression of Cytokine-inducible kinase, comprising operating a preferred target or more candidate modulators of Cytokine-inducible kinase, and comprising contacting a preferred target segment of the nucleic acid encoding Cytokine-inducible kinase with one or more modulators that modulate the expression of Cytokine-inducible kinase, and candidate modulators of Cytokine-inducible kinase with one or more modulators that modulate the expression of Cytokine-inducible kinase in a sample using at least one of the primers or probe comprising the action associated with Cytokine-inducible kinase in a sample using at least one of the primers or probe comprising the above compound; and a method of treating an animal having a disease or condition associated with Cytokine-inducible kinase is inhibited. The antisense oligonuclecide is useful for inhibiting the expression of Cytokine-inducible kinase is inhibited. The antisense oligonuclecide is useful for inhibiting the expression of expression of expression or prophylactic amount of the compound so that expression of Cytokine-inducible kinase is inhibited. The antisense oligonuclecide is useful for inhibiting the expression of expression or prophylactic amount of the compound so that expression of cytokine-inducible kinase is inhibited. The antisense oligonuclecide is useful for inhibiting the action of the compound so the expression of cytokine-inducible kinase is inhibited. The antisense oligonuclecide is useful for inhibiting the expr
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              treat diseases associated with the kinase expression, such as hyperproliferative disorders. In addition, the compound is used for disquostics, prophylaxis, or as research reagents or kits. This sequence represents a human cyctokine-inducible kinase antisense oligonucleotide.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           New antisense oligonucleotides useful for modulating Cytokine-inducible kinase expression, useful for diagnosing, preventing or treating conditions associated with aberrant kinase expression e.g. hyperproliferative disorders.
/note= "OTHER= Phosphorothioate backbone. All cytidines are 5-methylcytidines"
                                                                                                                                                       /*tag= a
/mod_base= OTHER
/note= "OTHER= 2'-0-Methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                       /mod_base= OTHER
/note= "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 20 BP; 4 A; 7 C; 6 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Example 15; SEQ ID NO 45; 56pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              2387 CATCCAGAGCCGCTGGTGT 2405
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         23-NOV-2002; 2002US-00304116.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 23-NOV-2002; 2002US-00304116.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.4%;
                                                                                                                                                                                                                                                                                                                                                     υ
                                                                                                                                                                                                                                                                                     15. .20
/*tag= c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Local Similarity oz...
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2004-399685/37.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Ward DT, Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    US2004101857-A1.
                                                                                                           modified base
                                                                                                                                                                                                                                                                                               modified base
```

N

```
cytostatic; gene therapy; breast cancer; polymorphism detection; PCR;
                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADQ29449
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT
   δ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                  The present invention relates to diagnosing or monitoring transplant rejection, e.g. cardiac or kidney transplant rejection, in an individual comprises detecting the expression level of one or more genes. The methods, system and kits are useful in diagnosing or monitoring transplant rejection, e.g. heart, kidney, liver, pancreas, pancreatic islet, lung, bone marrow or stem cell transplant rejection, e.g. heatt, kidney, liver, pancreas, pancreatic islet, lung, bone marrow or stem cell transplant rejection, in an individual. The method is also useful in assessing the immune status of an individual. The methods are also useful in diagnosing and monitoring diseases that involve the immune system, e.g. rheumatorid arthritis, lupus, inflammatory bowel diseases, multiple sclerosis, HIV/AIDS or viral, bacterial or fungal infection. The present sequence represents a primer for a 50 mer oligonucleotide marker for diagnosis and monitoring of allograft rejection and other disorders.
                                                                                                                                                                                                                                                                                                                                                    Diagnosing or monitoring transplant rejection, e.g. heart, kidney, liver, pancreas, pancreatic islet, lung, bone marrow or stem cell transplant rejection, in an individual, comprises detecting the expression level of
                                                                                                       transplant rejection; immune system; rheumatoid arthritis; lupus; inflammatory bowel disease; multiple sclerosis; HIV; AIDS; ss; primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                            Wohlgemuth J, Fry K, Woodward R, Ly N, Prentice J, Morris M;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Breast cancer associated polymorphism detection primer #5.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 20 BP; 5 A; 5 C; 6 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DB 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Score 15.8; DB 1
Pred. No. 2e+02;
                                                                                  Set 1 right PCR primer for marker probe #299
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                              Claim 58; SEQ ID NO 1294; 1762pp; English.
                                                                                                                                                                                                                                                                       EXPR-) EXPRESSION DIAGNOSTICS INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AGCAGGGCATGCCTGTGCT 2026
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AGCAGGAAATGCCTGTGCT 19
              BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ВР.
                                                                                                                                                                                                               24-APR-2003; 2003WO-US012946.
                                                                                                                                                                                                                                   24-APR-2002; 2002US-00131831.
20-DEC-2002; 2002US-00325899.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     0.4%;
              ADP11285 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                            (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADQ88907 standard; DNA;
                                                                                                                                                                                                                                                                                                                                WPI; 2004-400724/37.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Local Similarity
                                                                                                                                                                  WO2004042346-A2.
                                                                                                                                            Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              23-SEP-2004
                                                                                                                                                                                                                                                                                                         Rosenberg S;
                                                           12-AUG-2004
                                                                                                                                                                                        21-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            17;
                                                                                                                                                                                                                                                                                                                                                                                          the genes
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Н
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ADQ88907;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2008
                                    ADP11285;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RESULT 195
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          fatches
   ADP11285
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ADQ8890'
             ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  SEXEXEX
```

```
Identifying a subject at risk of breast cancer comprises detecting the presence or absence of one or more polymorphic variations associated with breast cancer in a nucleic acid sample from a subject.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            The invention describes a method of identifying (MI) a subject at risk of breast cancer. The method comprises detecting the presence or absence of one or more polymorphic variations associated with breast cancer in a nucleic acid sample from a subject. The methods, nucleic acids, proteins, and compositions are useful for diagnosing, preventing, and treating breast cancer. Also described is a method useful for identifying candidate therapeutics for treating breast cancer. This sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       antimicrobial; antidiabetic; antirheumatic; antiarthritic; gastrointestinal; antiinflammatory; neuroprotective; dermatological; virucide; hepatotropic; human; TNR-alpha; tumour necrosis factor alpha; survivin; TNR-alpha associated disorder; infection; diabetes; rheumatoid arthritis; Crohn's disease; pancreatitis; multiple sclerosis; atopic dermatitis; hepatitis; antisense oligonucleotide;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                represents a primer used to analyse polymorphisms associated with breast
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%; Score 15.8; DB 1; Length 20;
89.5%; Pred. No. 2e+02;
iive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 В.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Reneland
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Human TNF alpha antisense oligonucleotide seqid 380.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Seguence 20 BP; 7 A; 3 C; 5 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Kammerer SM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Example 2; Page 59; 83pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     2233 ATGAGCTAGTAAAGAATTT 2251
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1 ATGAGCCAGTACAGAATTT 19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Braun A,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          29-AUG-2003; 2003US-00652795.
                                                                                                                                                                                                                                                                  25-NOV-2003; 2003WO-US037831.
                                                                                                                                                                                                                                                                                                                                       25-NOV-2002; 2002US-0429136P.
24-JUL-2003; 2003US-0490234P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ADQ29449 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             antigense technology; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                (SEQU-) SEQUENOM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Roth RB, Nelson MR,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2004-517424/49.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   US2004142346-A1
                                                                                                                               WO2004055196-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens
                                                             Ното варіеля
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       07-OCT-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     22-JUL-2004.
                                                                                                                                                                                                    01-JUL-2004
primer; ss.
```

.. 0

10001863-3.81.rng

```
The invention relates to treating an inflammatory disorder in an individual comprising administering an oligonucleotide (an antisense cligonucleotide) up to 30 mucleotides in length complementary to a nucleic acid molecule encoding human tumour necrosis factor-alpha (TNF-alpha). The oligonucleotide useful in treating an inflammatory disorder inhibits the expression of the human tumour necrosis factor-alpha (TNF-alpha). The oligonucleotide usequences, given in the specification. The antisense oligonucleotide is equences, given in the specification. The antisense oligonucleotide is comprises at least one modified intersugar linkage. The intersugar linkage is a phosphorothioate linkage. The oligonucleotide further comprises at least one 2'-O-methoxyethyl modification and at least one 5'-methyl cytidine, where every 2'-O-methoxyethyl modified cytidine residue is a 5-methyl cytidine, and where every cytidine residue is a 5-methyl cytidine, and where every cytidine residue is a 5-methyl cytidine. The modified intersugar linkage is a methylene (methylimino)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             intersugar linkage. The methods and compositions of the present invention are useful for the diagnosis, prevention and/or treatment of diseases or conditions associated with aberrant expression or activity of the TNF-alpha, such as inflammarcry, infectious and autoimmune diseases, including diabetes, rheumatoid arthritis, Crohn's disease, pancreatitis,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Treating inflammatory disorders, such as diabetes, rheumatoid arthritis and multiple sclerosis, using antisense oligonucleotides targeted to nucleic acids encoding human tumor necrosis factor-alpha (TNF-alpha).
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       including diabetes, rheumatoid arthritis, Crohn's disease, pancreatitis, multiple sclerosis, atopic dermatitis, inflammatory bowel disease, colitis and hepatitis. The present sequence is an antisense
                           16. .20
/*tea= c
/mod base= OTHER
/note= "Phosphorothioate linkages and 2'-methoxyethyl
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0.4%; Score 15.8; DB 1; Length 20;
39.5%; Pred. No. 2e+02;
ive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      targeting the human TNFalpha gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Sequence 20 BP; 7 A; 4 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Shanahan WR;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Example 24; SEQ ID NO 380; 145pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Butler MM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        3414 TTTCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AAZ11675/c
ID AAZ11675 standard; DNA; 21 BP.
                                                                                                                                                                                                                                                                                            05-OCT-1998; 98US-00166186.
18-MAY-1999; 99US-00313932.
02-APR-2001; 2001US-00824322.
                                                                                                                                                                                                                                                     26-AUG-2003; 2003US-00647918.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     89.58;
         residues"
                                                                                                                   residues"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Baker BF, Bennett CF,
                                                                                                                                                                                                                                                                                                                                                                                  (BENN/) BENNETT C F.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 2004-580193/56.
                                                                                                                                                                                                                                                                                                                                                                                                                              (BUTL/) BUTLER M M. (SHAN/) SHANAHAN W
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Similarity
                                                                                                                                                             US2004152652-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        oligonucleotide
                           modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AA211675
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Local
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Best Loca
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     RESULT 198
         ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                  The invention describes a double stranded RNA compound 8-80 nucleobases
in length targeted to a nucleic acid molecule encoding human TNF-alpha,
where the compound specifically hybridises with the nucleic acid molecule
encoding TNF-alpha and inhibits the expression of survivin. Also
described is a double stranded RNA compound having a fully defined
sequence of 20 bp (SEQ ID NO: 432) as given in the specification. Also
disclosed are TNF-alpha polypeptides, host cells, vectors and antibodies
used in the methods of the invention. The methods and compositions of the
present invention are useful for the diagnosis, prevention and/or
treatment of diseases or conditions associated with aberrant expression
treatment of disease, pancreatitis, multiple sclerosis, atopic
dermatitis and hepatitis. This sequence represents a human tumour
ecrosis factor alpha (TNF-alpha) antisense oligonucleotide.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                     new notable stranded RNA compound inhibiting expression of human TNF-alpha and survivin, useful for diagnosing, preventing or treating infection, diabetes, arthritis, multiple sclerosis and hepatitis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Human; tumour necrosis factor alpha; TNFalpha; ss; antisense gene therapy; inflammatory disorder; phosphorothioate linkage; methylene(methylimino) intersugar linkage; infection; autoimmune disease; diabetes; rheumatoid arthritis; Crohn's disease; pancreatitis; multiple solerosis; atopic dermatitis; inflammatory bowel disease; colitis; hepatitis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /*tag= a
/mod_base= OTHER
/note= "Phosphorothioate linkages and 2'-methoxyethyl
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Antisense oligonucleotide targeting human TNFalpha ISIS104785.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       *tag= b
mod_base= OTHER
'note= "All Cytidines are 5-merthylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Score 15.8; DB 1; Length 20;
Pred. No. 2e+02;
0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 7 A; 4 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                       Shanahan WR;
                                                                                                                                                                                                                                                                                                                                                                                  Example 24; SEQ ID NO 380; 156pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Location/Qualifiers
                                                                                                                                                                                                       Butler MM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      3414 TTTCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1 rcrcaaggaagrcrggaaa 19
05-OCT-1998; 98US-00166186.
18-MAY-1999; 99US-00313932.
02-APR-2001; 2001US-00824322.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Local Similarity 89.5%;
les 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADR02965 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (first entry)
                                                                                                                                                                                                  Baker BF, Bennett CF,
                                                                                                                                 (BUTL/) BUTLER M M.
(SHAN/) SHANAHAN W R.
                                                                                     (BENN/) BENNETT C F.
                                                                                                                                                                                                                                                WPI; 2004-552557/53.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     21-OCT-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ADR02965
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 197
```

ò g

10001863-3.sl.rng

```
Synthetic.
Mus sp.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AAS22336;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 200
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AAS22336
  ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                       The invention provides methods for identifying an Epstein Barr Virus (EBW) infection, that comprises detarmining viral gene transcription patterns by amplification of specific RNA sequences. The binding sites of the oligos suitable for amplification are located in the following genes: Epstein Barr early RNA (EBER-1), Epstein Barr nuclear antigen 1 (EBNA-1). Elatent membrane protein 1 (LMP-1), LMP-2, and vILIO (BCRF-1), BARF1 and BDLF2. The method comprises (a) amplifying a target sequence within one or more RNA(s) transcribed from above gene sequences and the (b) and identifying the amplified products, determining the transcription pattern and identifying a transcription based amplification technique such as amplified using a transcription based amplification technique such as succiated diseases. Sequences AAZI1672-75 represent oligos specific for BARF-1 RNA. (Updated on 27-AUG-2003 to correct OS field.)
                                                       Epstein Barr Virus; EBV infection; viral; gene transcription; EBER-1; Epstein Barr early RMs; Epstein Barr nuclear antigen 1; EBNA-1; LMP-1; latent membrane protein; LMP-2; VILLO; BCRF-1; BARF1; BDLF2; NASBA; EBV-associated malignancy; primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                MSP; macrophage stimulating protein; apoptosis; murine; treatment; neuroendocrine cell; RON receptor; small cell lung carcinoma; tumour; pathogen infection; thrombocyte production; megakaryocyte maturation; thrombocytopaenia; hepatocyte growth; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.4%; Score 15.8; DB 1; Length 21; 89.5%; Pred. No. 2.2e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               2; Indels
                                                                                                                                                                                                                                                                                Vervoort MBHJ, Van Den Brule AJC, Middeldorp JM;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 21 BP; 5 A; 3 C; 9 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0; Mismatches
                                                                                                                                                                                                                                                                                                                             Identifying Epstein Barr Virus infection
                                   Oligo specific for EBV BARF-1 RNA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        2319 GCTGCCAACATCATCCATG 2337
                                                                                                                                                                                                                                                                                                                                                     Claim 11; Page 21; 50pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             gergeckáckrerecekre 3
                                                                                                                                                                                                 99WO-EP001392.
                                                                                                                                                                                                                       98EP-00200655,
98EP-00204231,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AAV72092 standard; cDNA; 21
  (revised)
(first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    12-APR-1999 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
Best Local Similarity 89.5'
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Mouse MSP PCR primer #2.
                                                                                                                                                                                                                                                          (ALKU ) AKZO NOBEL NV.
                                                                                                                             Human herpesvirus 4.
                                                                                                                                                                                                                                                                                                       WPI; 1999-551051/46.
                                                                                                                                                    WO9945155-A2
                                                                                                                                                                                                 01-MAR-1999;
                                                                                                                                                                                                                                    14-DEC-1998;
                                                                                                                                                                                                                        04-MAR-1998;
 27-AUG-2003
19-NOV-1999
                                                                                                                                                                           10-SEP-1999
                                                                                                                  Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AAV72092;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               21
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 199
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             셤
```

```
AAV72085-V72099 represent PCR primers and probes used in the isolation and amplification of novel human and murine macrophage stimulating protein, MSP, which are used in a method for the prophylactic treatment of a tumour derived from neuroendocrine cells (NEC) by administration of this MSP to a subject at risk, sufficient to induce apoptosis of NEC expressing a RON receptor (the receptor for MSP). The method is used to treat or prevent small cell lung carcinoma and apoptosis of RON-expressing cells may be induced in vivo or in vitro. Screening NEC from subject for susceptibility to MSP-induced apoptosis is used to identify patients who will benefit from treatment with the MSP protein. MSP is already known for treating pathogen infections, for stimulating thrombocytopaenia) and for stimulating growth of cells (particularly
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Human; collagen; COLIA1; COLIA2; COL9A1; COL9A2; COL9A3; se; osteoporosis; multiple epiphyseal dysplasia; osteogenesis imperfecta; shortness of stature; low bone density; gene therapy; PCR primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                    Treating tumours derived from neuroendocrine cells with macrophage stimulating protein - or its nucleic acid, also for preventing development of these tumours, specifically small cell lung carcinoma.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%; Score 15.8; DB 1; Length 21; 89.5%; Pred. No. 2.2e+02; ive 0; Mismatches 2; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 21 BP; 4 A; 9 C; 2 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Human COL9A2 PCR primer 1 for Exon 29.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Example 2; Page 71; 100pp; English
                                                                                                                                                                                                                                                            (BGHM ) BRIGHAM & WOMENS HOSPITAL.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2634 GAAGGAACAGTGGGTACAG 2652
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     N
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAS22336 standard; DNA; 21 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20 GAAGGAGCAGTGGGTACTG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 97US-00943731,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            91US-00803628.
94US-00212322.
                                                                                                                         98WO-US011573.
                                                                                                                                                                                              97US-0048594P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        17; Conservative
                                                                                                                                                                                                                                                                                                                              Sunday ME, Willet C;
                                                                                                                                                                                                                                                                                                                                                                                            WPI; 1999-059877/05.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens
                                                                                                                                04-JUN-1998;
                                                                                                                                                                                              04-JUN-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               24-OCT-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 US6265157-B1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 03-OCT-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            03-DEC-1991;
13-MAR-1994;
WO9855141-A1
                                                          10-DEC-1998.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             hepatocytes)
```

. 0 10001863-3.sl.rng

2001US-0272414P. 2001US-0272787P. 2001US-0272922P.

2001US-0272405P 2001US-0272410P

2001US-0273300P.

2001US-0273048P 2001US-0276401P 2001US-0277324P

2001US-0278660P. 2001US-0280039P. 2001US-0280234P.

2001US-0283443P. 2001US-0285754P. 2001US-0286096P.

2001US-0280818P

2001US-0288353P. 2001US-0291703P.

2001US-0299695P 001US-0303242P 2001US-0312858P. 2001US-0313280P. 2001US-0315614P.

```
Human; NOVX; atherosclerosis; hypertension; obesity; cancer; cytostatic; hypotensive; antiarteriosclerotic; anorectic; gene therapy; NOV72; PCR;
                                                                                                                                                                                                                                                                    0; Mismatches
                                                                                                                                                                                                                                                                              1921 AACACCAGAGTTTCCTGCA 1939
                                                                                                                                                                                                                                                                                    AACACCAGAGTCTCCTCCA 19
                                                                                  Calim 8; Fig24; 617pp; English
                                                                                                                                                                                                                                                                                                                                                     NOV72 PCR primer, SEQ ID 476.
                                                                                                                                                                                                                                                         Match 0.4%;
Local Similarity 89.5%;
                                                                                                                                                                                                                                                                                                                    ADH49192 standard; DNA; 21
                                                                                                                                                                                                                                                                                                                                          25-MAR-2004 (first entry)
                                                                                                                                                                                                                                                                   17; Conservative
                                                  WPI; 2001-432201/46.
                                Westerĥausen
Ala-Kokko L,
                                                                                                                                                                                                                                                                                                                                                                                       Homo sapiens
                           Prockop DJ,
                                                                                                                                                                                                                                                                                                                                                                            primer; 88.
                                                                                                                                                                                                                                                                                         ٦
                                                                                                                                                                                                                                                                                                                                ADH49192;
                                                                                                                                                                                                                                                         Query Match
                                                                                                                                                                                                                                                                                                          Matches
ઢ
                                                                                                                                                                                                                                                                                       g
```

```
26-FEB-2002; 2002WO-US005910
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (CURA-) CURAGEN CORP.
                                                                                                                                                                                                                                             02-MAR-2001;
16-MAR-2001;
20-MAR-2001;
                                                                                                                                                                                02-MAR-2001;
02-MAR-2001;
02-MAR-2001;
                                                                                                                                                                                                                                                                                                                              30-MAR-2001;
30-MAR-2001;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     16-AUG-2001;
                                                                                                                                                               28-FEB-2001;
                                                                                                                                                                                                                                                                                                           20-MAR-2001;
                                                                                                                                                                                                                                                                                                                                                                       02-APR-2001;
                                                                                                                                                                                                                                                                                                                                                                                               12-APR-2001;
                                                                                                                                                                                                                                                                                                                                                                                                              23-APR-2001;
                                                                                                                                                                                                                                                                                                                                                                                                                                    24-APR-2001;
                                                                                                                                                                                                                                                                                                                                                                                                                                                       03-MAY-2001;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               17-MAY-2001;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                31-MAY-2001;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           21-JUN-2001;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    13-AUG-2001;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      20-JUN-2001;
The invention relates to Detecting a collagen gene alteration associated with a pathological condition in a human subject by obtaining from the subject a sample nucleic acid containing a portion of at least 15 consecutive nucleotides of the segment of the COLIA1 gene extending in the 5' to 3' direction from 78 nucleotides of intron 27 located adjacent consecutive nucleotide and a first and second site, obstrion contains an intronic nucleotide and a first and second site, determining the sequence of the portion and comparing the sequence of the portion with the corresponding consensus sequence of the COLIA1 gene where a difference between the sequence of the portion and the consensus sequence in the presence of the portion and the consensus sequence indicates the presence of the collagen alteration in the subject. The method is used for detecting abnormalities in a COLI or COL9 gene is useful for determining whether a subject is afflicted with pathological conditions associated with an altered collagen gene such as osteoporosis, multiple companies of the collagen gene such as osteoporosis, multiple companies of the consensus of stature and companies of section of an abnormality in a collagen gene is also useful for designing a therapeutic nucleotide or gene therapy agent conding and non-coding sequences of any of the COLIA or COL9 genes.

Therefore the method can be used to detect collagen gene alterations in both the coling and non-coding sequences of any of the COLIA or COL9 genes.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                of
                                                                                                                                                                                                                       Detecting collagen gene alteration, useful for diagnosing osteoporosis, multiple epiphyseal dysplasia, osteogenesis imperfecta, shortness of stature and low bone density in humans.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        which affect either the primary sequence of a collagen protein chain, splicing of the mRNA encoding such chains or regulation of expression
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   the genes encoding such chains. The present sequence is a PCR primer which amplifies a nucleic acid from a collagen gene of the invention
                                                                                                                    Koerkkoe J;
Paassilta P;
                                                                                            Spotila LD, Deltas CD, Sereda L;
Larson A, Pack M, Colige A, Early J,
Annunen S, Pihlajamaa T, Vuoristo M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence 21 BP; 6 A; 8 C; 3 G; 4 T; 0 U; 0 Other;
        (UYAL-) UNIV ALLEGHENY HEALTH SCI.
(UYJE-) UNIV JEFFERSON THOMAS.
(UYOU-) UNIV OULU.
            (UYAL-)
```

Alsobrook JP, Anderson DW, Ballinger RA, Boldog FL, Burgess CE; Casman SJ, Ellerman KE, Gangolli EA, Gerlach VL, Gilbert JA, Gorman L, Guo X, Gueev VY, Kekuda R, Li L, Liu X, Malyankar UM, Miller CE, Millet I, Padigaru M, Patturajan M, Pena CEA, Peyman J Rastelli L, Shenoy SG, Shimkets RA, Smithson G, Spytek KA, Stone Taupier RJ, Tchernev VT, Vernet CAM, Zerhusen BD; New NOVX polypeptides or polynuclectides, useful for preventing or treating disorders or syndromes e.g., atherosclerosis, hypertension. WPI; 2002-698672/75. ö Gaps ö Score 15.8; DB 1; Length 21; Pred. No. 2.2e+02; 2; Indels

obesity or cancer.

Peyman JA; , Stone DJ;

The present invention relates to novel human NOVX proteins, where X is any number from 1 to 91 and their coding sequences (see ADH48717-ADH48930). The proteins and coding sequences are useful for preventing or treating disorders or syndromes e.g. atherosclerosis, hypertension, obesity or cancer. The present sequence was used in an example from the invention. Example 2; Page 855; 923pp; English.

Match 0.4%; Score 15.8; DB 1; Length 21; Local Similarity 89.5%; Pred. No. 2.2e+02; Bes 17; Conservative 0; Mismatches 2; Indels 2626 GGAATCCAGAAGGAACAGT 2644 Matches .17; Conservative Query Match

ð

Sequence 21 BP; 2 A; 5 C; 5 G; 9 T; 0 U; 0 Other;

ö

Gaps

ö

m GGAAACCACAAGGAACAGT 7

AAQ90158 standard; DNA; 20 BP RESULT 202 AAQ90158/c ID AAQ901

WO200268652-A2

06-SEP-2002

```
Stinchcomb DT,
                                                                                                                                                                                                                              and in cancers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WO200058519-A2
      18-MAY-1994;
13-JAN-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       31-MAR-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Altshuler D,
Lipshutz RJ,
                                                                                                                                                                                                                                                                                                                                                                                             09-FEB-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    05-OCT-2000.
                                                                                                                                                                                                                                                                                                                                                                             AAC70462;
                                                                                                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                           RESULT 204
                                                                                                                                                                                                                                                                              Matches
                                                                                                                                                                                                                                                                                                                                                    AAC70462
ò
                                                                                                                                                                                                                                                                                                                qq
                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                       The primers given in AAQ90157-58 are based on a portion of an allergen purified from Japanese cedar pollen, and were used to isolate clone SC09 bearing a partial sequence (nt 173-240 of the sequence given in AAQ90156) of the allergen gene by PCR amplification of pollen-derived cDNA. (Updated on 25-WAR-2003 to correct PN field.)
                                                                                                                                                                                                                                                               New Japanese cedar pollen allergen polypeptide - and DNA coding for it, useful for treatment and diagnosis of cedar pollen allergy.
                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human c-myb hammerhead ribozyme target sequence (nt. position 2889)
                                                                  Japanese cedar; pollen; allergen; allergy; therapy; diagnostic; desensitizer; Cryptomeria japonica; polymerase chain reaction; PCR;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Enzymatic nucleic acid; hammerhead; ribozyme; cleavage; human; smooth muscle cell; hyperproliferation; restenosis; cancer; c-myb; coronary angioplasty; ss.
                                                                                                                                                                                                                                                                                                                                                                                               .
0
                                                                                                                                                                                                                                                                                                                                                                             ch 0.4%; Score 15.6; DB 1; Length 20; l Similarity 83.3%; Pred. No. 2.1e+02; 15; Conservative 2; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                            Sequence 20 BP; 2 A; 5 C; 6 G; 5 T; 0 U; 2 Other;
                                                                                                                                                                                                            (HAYB ) HAYASHIBARA SEIBUTSU KAGAKU.
                                                                                                                                                                                                                                                                                         Disclosure, Page 8; 41pp; English.
                                                                                                                                                                                                                              Kurimoto M;
                                                                                                                                                                                                                                                                                                                                                                                                                2660 TTGGCAGGAAGCAACATC 2677
                                                                                                                                                                                                                                                                                                                                                                                                                          BP.
                                                 Pollen allergen gene primer 2.
                                                                                                                                                                           93JP-00299151.
                                                                                                                                                                                  93JP-00344596.
93JP-00346814.
                                                                                                                                                          94EP-00308117.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     95WO-US006368
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAT81553 standard; RNA; 17
                        (revised)
(first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               14-DEC-1997 (first entry)
                                                                                                                                                                                                                              Namba M, Torigoe K,
                                                                                                                                                                                                                                               WPI; 1995-195588/26
                                                                                                                                                                                                                                                                                                                                                                                       Best Local Similarity
                                                                                                                                                                         05-NOV-1993;
20-DEC-1993;
27-DEC-1993;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     18-MAY-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO9531541-A2
                                                                                                                                                          03-NOV-1994;
                                                                                                                       EP655500-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     23-NOV-1995
                       25-MAR-2003
01-NOV-1995
                                                                                                                                       31-MAY-1995
                                                                                      primer; ss.
                                                                                                      Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AAT81553;
        AAQ90158;
                                                                                                                                                                                                                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                                                               Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                          à
                                                                                                                                                                                                                                                                                                                                                                                                                               g
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   enzymatic nucleic acid, especially a harmerhead ribozyme, which cleaves the human c-myb sequence at the base position indicated in the descriptor line. The c-myb sequence was screened for optimal ribozyme target sites using a computer folding algorithm, and regions of the mRNA which did not form secondary folding structures and contained potential ribozyme cleavage sites were identified. Ribozymes were synthesised and their activities optimised by either varying the length of the binding arms or by modification to prevent degradation by nucleases. The ribozymes cleave the c-myb sequence and can be used to prevent smooth muscle cell hyperproliferation in restenosis, especially after coronary angioplasty,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Single nucleotide polymorphism; SNP; human; genetic disease; disease susceptibility; cardiovascular system; endocrine system; neurological system; forensic testing; paternity testing; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                   New enzymatic nucleic acid molecules - cleave RNA produced by e.g. c-myb, for treating restenosis or cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Nucleic acid selected from one of 106 genes comprising single nucleotide polymorphisms, allele-specific oligonucleotides to the genes are useful
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            The present sequence represents the preferred target sequence for an
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Lander ES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Match 0.4%; Score 15.4; DB 1; Length 17; Local Similarity 23.5%; Pred. No. 1.7e+02; Local 4; Conservative 12; Mismatches 1; Indels
                                                                                                                                                                              Jarvis T;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 17 BP; 5 A; 0 C; 0 G; 0 T; 12 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Daley GQ, Ireland JS, Sklar P;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Single nucleotide polymorphism PCR primer #195.
                                                                                                                                                                                  Draper K, Mcswiggen J,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (WHED ) WHITEHEAD INST BIOMEDICAL RES
                                                                                                                                                                                                                                                                                                                                                                                                                                        Claim 1; Page 78; 128pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           3305 TTTTTTTTTTTTT 3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1 UUUUUAAUUUUUAUAUAU 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ВЪ
94US-00245466.
95US-00373124.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               30-MAR-2000; 2000WO-US008440.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       99US-0127248P
                                                                                                          (RIBO-) RIBOZYME PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AAC70462 standard; DNA; 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Cargill M,
Patil N, S
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (AFFY-) AFFYMETRIX INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2000-611722/58.
                                                                                                                                                                                                                                                          WPI; 1996-010927/01.
```

ö

Gaps

ö

(first entry)

Length 17; Indels

```
Claim 8; Fig 5; 214pp; English.
              8; Fig 5; 214pp; English
                                                                                                               1 ACAGGAAGCCCCATCCA 17
                                                                                                                                            BP.
                                                                                     0.4%;
94.1%;
                                                                                                                                                                                                                                     30-MAR-2000; 2000WO-US008440
                                                                                                                                                                                                                                              99US-0127248P
                                                                                                        492 ACAGGAAACCCCATCCA
                                                                                                                                           AAC70489 standard; DNA; 17
                                                                                              16; Conservative
                                                                                                                                                                                                                                                                       Cargill M,
Patil N, 8
                                                                                                                                                                                                                                                             (AFFY-) AFFYMETRIX INC
                                                                                                                                                                                                                                                                                      WPI; 2000-611722/58.
                                                                                    Query Match
Best Local Similarity
     genetic analysis.
                                                                                                                                                                                                                                                                                                               genetic analysis.
                                                                                                                                                                                                                WO200058519-A2
                                                                                                                                                                                                                                                                       Altshuler D,
Lipshutz RJ,
                                                                                                                                                               09-FEB-2001
                                                                                                                                                                                                       Homo sapiens
                                                                                                                                                                                                                                               31-MAR-1999;
                                                                                                                                                                                                                          05-OCT-2000.
                                                                                                                                                     AAC70489;
                                                                diseases
              Claim
                                                                                                                                  RESULT 205
                                                                                               Matches
                                                                                                                                      ઠે
                                                                                                                a
```

```
virucide; neuroprotective; antibacterial; replication; pancreatitis; encephalitis; myocarditis; meningitis; infection; hepatitis; liver failure; cancer; cirrhosis; Hammerhead; Inozyme; DNAzyme;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WNV; West Nile Virus; antiinflammatory; cytostatic; hepatotropic;
                                                                                    Query Match 0.4%; Score 15.4; DB 1; I
Best Local Similarity '94.1%; Pred. No. 1.7e+02;
Matches 16; Conservative 0; Mismatches 1;
                                               Sequence 17 BP; 6 A; 7 C; 3 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WNV Inozyme substrate SEQ ID NO 3134.
                                                                                                                                                                            492 ACAGGAAACCCCATCCA 508
                                                                                                                                                                                                                   1 ACAGGAAGCCCCATCCA 17
                                                                                                                                                                                                                                                                                                                                  ACN03131 standard; RNA; 17 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           19-OCT-2001; 2001WO-US048350.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      20-OCT-2000; 2000US-0242411P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Blatt L, Mcswiggen JA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Amberzyme; Zinzyme; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (RIBO-) RIBOZYME PHARM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            MCSW/) MCSWIGGEN J A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         West Nile Virus.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WO200268637-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     BLATT
                                                                                                                                                                                                                                                                                                                                                                                                                        22-APR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  06-SEP-2002.
                                                                                                                                                                                                                                                                                                                                                                             ACN03131;
      diseases
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     BLAT/)
                                                                                                                                                                                                                                                                                       RESULT 206
  SXS
                                                                                                                                                                            ઠ
                                                                                                                                                                                                                   셤
                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                   The present invention is concerned with a number of human single nucleotide polymorphisms (SNPs) which the inventors identified in human genes. These SNPs can be used in disease diagnosis and prediction of an individual's susceptibility to disease, in forensic and paternity testing and in genetic mapping. In particular, the SNPs of the invention can be used to diagnose susceptibility to diseases of the cardiovascular, endocrine and neurological systems, such as coronary artery disease, echizophrenia, cancer, autoimmune diseases, Alzheimer's and Parkinson's
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  The present invention is concerned with a number of human single nucleotide polymorphisms (SNPs) which the inventors identified in human genes. These SNPs can be used in disease diagnosis and prediction of an individual's susceptibility to disease, in forensic and paternity testing and in genetic mapping. In particular, the SNPs of the invention can be used to diagnose susceptibility to diseases of the cardiovascular, endocrine and neurological systems, such as coronary artery disease, schizophrenia, cancer, autoimmune diseases, Alzheimer's and Parkinson's
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      disease susceptibility; cardiovascular system; endocrine system;
neurological system; forensic testing; paternity testing; PCR primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Nucleic acid selected from one of 106 genes comprising single nucleotide polymorphisms, allele-specific oligonucleotides to the genes are useful for phenotypic correlations, forensics, paternity testing, medicine and
for phenotypic correlations, forensics, paternity testing, medicine and
                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Single nucleotide polymorphism; SNP; human; genetic disease;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          , Daley GQ, Ireland JS, Lander ES;
Sklar P;
                                                                                                                                                                                                                                                                                                                                                                             Length 17;
                                                                                                                                                                                                                                                                                                                                                                                                                      1; Indels
                                                                                                                                                                                                                                                                                                                                Sequence 17 BP; 6 A; 7 C; 3 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                         Score 15.4; DB 1;
Pred. No. 1.7e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Single nucleotide polymorphism PCR primer #213.
                                                                                                                                                                                                                                                                                                                                                                                                                        0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (WHED ) WHITEHEAD INST BIOMEDICAL RES
```

```
..
0
                                                                                                                                                                                                                                                                                                                                                                                                  The invention relates to nucleic acid molecules that modulate replication of the West Nile Virus (WNV). The nucleic acid molecules are useful for treating a condition related to WNV infection e.g. pancreatitis, encephalitis, myozaditis, meningitis, neurologic infection, hepaticis, liver failure, hepatocellular carcinoma or cirrhosis. The nucleic acid molecule is selected from the group of ribozymes consisting of Hammerhead, Inozyme, G-cleaver, DNAzyme, Amberzyme and Zinzyme. The nucleic acid molecules further comprise at least five ribose residues, at least ten 2'-O-methyl modifications, phosphorothicate linkages on at least three of the 5' terminal nucleotides and a 3' end modification of a 3'-3' inverted abasic moiety. Nucleic acid molecules SEQ ID No 1 to 37080 are claimed, however, SEQ ID No 2194-2206 and 17502-17514 are not given in the specification. The present sequence is that of a nucleic acid
                                                                                                New nucleic acid molecule that modulates replication of West Nile Virus (WNV), useful for treating a condition related to WNV infection e.g. pancreatitis, meningitis, hepatocellular carcinoma or cirrhosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match 0.4%; Score 15.4; DB 1; Length 17; Best Local Similarity 82.4%; Pred. No. 1.7e+02; Matches 14; Conservative 2; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 17 BP; 4 A; 4 C; 7 G; 0 T; 2 U; 0 Other;
                                                                                                                                                                                                                                                                                                       Claim 23; SEQ ID NO 3134; 495pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                molecule of the invention
WPI; 2002-706994/76.
```

Tumour suppression related human fukutin oligo SEQ ID No 1236,

(first entry)

12-JUN-2003

ABT35599;

ABT35599 standard; DNA; 17 BP.

ABT35599/c

```
The invention relates to nucleic acid molecules that modulate replication of the West Nile Virus (WNV). The nucleic acid molecules are useful for treating a condition related to WNV infection e.g. pancreatitis, encephalitis, myocarditis, meningitis, neurologic infection, hepatitis, liver failure, hepatocellular carcinoma or cirrhosis. The nucleic acid molecule is selected from the group of ribozymes consisting of Hammerhead, Inozyme, G-cleaver, DNAzyme, Amberzyme and Zinzyme. The nucleic acid molecules further comprise at least five ribose residues, least ten 2'-O-methyl modifications, phosphorothioate linkages on at least three of the 5' terminal nuclectides and a 3' end modification of a 3'-3' inverted abasic moiety. Nucleic acid molecules SEQ ID NO 1 to 37080 are claimed; however, SEQ ID NO 2194-2206 and 17502-17514 are not given in the specification. The present sequence is that of a nucleic acid
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         New nucleic acid molecule that modulates replication of West Nile Virus (WNV), useful for treating a condition related to WNV infection e.g. pancreatitis, meningitis, hepatocellular carcinoma or cirrhosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           WNV; West Nile Virus; antiinflammatory; cytostatic; hepatotropic; viruoids; neuroprotective; antibacterial; replication; panoreatitis; encephalitis; myocarditis; meningitis; infection; hepatitis; liver failure; cancer; cirrhosis; Hammerhead; Inozyme; DNAzyme;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WNV minus strand Inozyme substrate SEQ ID NO 9951.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 17 BP; 2 A; 7 C; 4 G; 0 T; 4 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Claim 23; SEQ ID NO 9951; 495pp; English
GACACAGATGGCTGGGA 3774
                                            |||||||| :|||:||||
GACACAGCUGGCUGGGA 17
                                                                                                                                                                                                                                                                           ВP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          19-OCT-2001; 2001WO-US048350
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20-OCT-2000; 2000US-0242411P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (RIBO-) RIBOZYME PHARM INC. (BLAT/) BLATT L.
                                                                                                                                                                                                                                                                       ACN09948 standard; RNA; 17
                                                                                                                                                                                                                                                                                                                                                                                                                     22-APR-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               molecule of the invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Blatt L, Mcswiggen JA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Amberzyme; Zinzyme; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (MCSW/) MCSWIGGEN J A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 2002-706994/76.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   West Nile Virus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WO200268637-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             06-SEP-2002.
                                                                         Н
                                                                                                                                                                                                                                                                                                                                                ACN09948;
3758
                                                                                                                                                                                      RESULT 207

ACN09948/C

DD 22-APR

XX ACN099

XX ACN099

XX ACN099

XX WANV, W

WANV, W

KW HIVOR

KW AMBERZ

XX AMBERZ

XX ACN099

XX HOLD

PP 19-OCT

XX AMBERZ

XX
ò
                                                                    d
```

```
The invention relates to a novel isolated 17 mer nucleic acid sequence, given in the specification, a sequence containing at least 15 consecutive uncleotides from the 17 mer sequence, a sequence with, after optimal alight and the specification, a sequence with, after optimal alight and primars for them the 17 mer sequence. The sequence that the single to them under highly stringent conditions, or the complement of any of them, or the corresponding RNA. The novel isolated nucleic cides of the invention are useful as probes and primers for detecting, dentifying and/or amplifying a nucleic acid, e.g. as one component of a gene chip, in vitro as (anti) sense reagents, and for production of recombinant polypeptides. Any of the nucleic acids, containing the nucleic acids, cells containing the nector or antibodies directed against the polypeptides are useful for preparation of pharmaceuticals for prevention and/or treatment of viral degeneration, specifically cancer but also Alzheimer's disease and containing the expression of the 17 meruleic acids in patient samples is useful for disposis and/or prognosis of these containing the polypeptides can also be used to generate antibodies, and chips. The nucleic acid sequences of the invention can be used in gene therapy. This polymucleotide sequence represents a tumour suppression containing the related human fukutin oligonucleotide of the invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                 Cytostatic; virucide; neuroprotective; nootropic; neuroleptic; gene chip; antisense; sense; tumour; cell degeneration; cancer; Alzheimer's disease; edizophrenia; protein chip; gene therapy; tumour suppression; human fukutin; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          New isolated nucleic acid, useful for treating viral diseases associated with tumors and cell degeneration, also related polypeptides, antibodies
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0.4%; Score 15.4; DB 1; Length 17; 94.1%; Pred. No. 1.7e+02; tive 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 17 BP; 6 A; 4 C; 1 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Disclosure; Page 177; 720pp; French.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Tuijnder M;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               2411 TGAATATGAGATTGCTC 2427
                                                                                                                                                                                                                                                                                                                                                                                                                                             (MOLE-) MOLECULAR ENGINES LAB
                                                                                                                                                                                                                                                                                                                                                                  17-SEP-2002; 2002WO-IB004208.
                                                                                                                                                                                                                                                                                                                                                                                                   17-SEP-2001; 2001FR-00011978
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    and transfected cells.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Amson R,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2003-313353/30.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Local Similarity
                                                                                                                                                                                                                                                                                        WO2003025175-A2.
                                                                                                                                                                                                                                                 Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Telerman A,
                                                                                                                                                                                                                                                                                                                            27-MAR-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     11
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT 209
ACD54465
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Matches
8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 셤
```

ACD54465 standard; RNA; 17 BP.

ū

ö

Gaps

.; 0

0.4%; Score 15.4; DB 1; Length 17; 94.1%; Pred. No. 1.7e+02; ive 0; Mismatches 1; Indels

Query Match
Best Local Similarity 94.1
Matches 16; Conservative

3758 GACACAGATGGCTGGGA 3774

17 GACACAGCTGGCTGGGA

RESULT 208

10001863-3.81.rng

Best Local Similarity

```
Nucleic acid molecule, Hepatitis C virus; HCV; Hepatitis B virus; HBV; RNA stability; RNA expression; RNA synthesis; antisense; enzymatic nucleic acid; hammerhead ribozyme; DNAzyme; inozyme; zinzyme;
                                                                                                                                                                  amberzyme; G-cleaver ribozyme; decoy molecule; aptamer;
HBV reverse transcriptase; Enhancer I region; viral replication;
degenerative; disease state; HBV infection; HCV infection; cirrhosis;
liver failure; hepatocellular carcinoma; hepatotropic; cytostatic;
virucide; antiinflammatory; substrate; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Mcswiggen J, Morrissey D, Pavco P,
                                               HBV DNAzyme substrate sequence #24
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        26-MAR-2001, 2001US-00817879.
08-JUN-2001, 2001US-00877478.
08-JUN-2001, 2001US-0296876P.
24-OCT-2001, 2001US-0335059P.
05-DEC-2001, 2001US-0337055P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                              26-MAR-2002; 2002WO-US009187.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RIBOZYME PHARM INC.
24-SEP-2003 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Macejak D,
Roberts E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       MACEJAK D.
MCSWIGGEN J.
MORRISSEY D.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           WPI; 2003-229207/22
                                                                                                                                                                                                                                                                                                                 Hepatitis B virus.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      LEE P.
DRAPER K.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ROBERTS E
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  PAVCO P.
                                                                                                                                                                                                                                                                                                                                                              WO200281494-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   BLATT
                                                                                                                                                                                                                                                                                                                                                                                                              17-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Blatt L, N
Draper K,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (PAVC/)
(LEEP/)
(DRAP/)
(ROBE/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RIBO-)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            MACE/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           MORR/)
```

treating cirrhosis, liver failure, or condition associated with hepatitis C virus Example 1; Page 184; 387pp; English compound useful for hepatocellular carcinoma, infection.

The present invention relates to nucleic acid molecules which modulate the synthesis, expression and/or stability of Hepatitis C virus (HCV) or Hepatitis B virus (HBV) RNA. The nucleic acid molecules include antisense and enzymatic nucleic acids such as hammerhead ribozymes, DNAzymes, amberzymes, anderzymes, anderzymes, anderzymes, anderzymes, anderzymes, anderzymes, and stability of cleaver ribozymes. Also disclosed are nucleic acid decoy molecules and aptamers that bind to HBV reverse transcriptase and/or HBV reverse transcriptase primer sequences, as well as oligonuclectides that specifically bind the Enhancer I region of HBV DNA. The nucleic acids may be used to modulate the expression of HBV COMPOUNDS and HBV repeased and HBV viral replication. Also disclosed is a method for screening compounds and/or potential therapies directed against HBV, and compounds that modulate the expression and/or replication of HCV. The compounds maddisate the expression and HCV infection, replication and gene expression such as cirrhosis, liver failure, and hepatocellular carcinoma. The present sequence represents a substrate for one of the HBV infectionme, inosyme, declaryme, inosyme, declaryme, zinzyme, DNAzyme or amberzyme sequences disclosed in the present invention

Sequence 17 BP; 4 A; 4 C; 3 G; 0 T; 6 U; 0 Other;

0.4%; Score 15.4; DB 1; Length 17;

Query Match

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              The present invention relates to nucleic acid molecules which modulate the synthesis, expression and/or stability of Hepatitis C virus (HCV) or Hepatitis B virus (HBV) RNA. The nucleic acid molecules include antisense and enzymatic nucleic acids such as hammerhead ribozymes, DNAzymes, inozymes, zinzymes, amberzymes, and G-cleaver ribozymes. Also disclosed are nucleic acid decoy molecules and aptemers that bind to HBV reverse transcriptase and/or HBV reverse transcriptase primer sequences, as well as oligonucleotides that specifically bind the Enhancer I region of HBV genes and HBV viral replication. Also disclosed is a method for screening compounds and/or potential therapies directed against HBV, and compounds
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Novel compound useful for treating cirrhosis, liver failure, hepatocellular carcinoma, or condition associated with hepatitis C virus
                                                                                                                                                                                                                                                                     Nucleic acid molecule, Hepatitis C virus, HCV; Hepatitis B virus, HBV; RNA stability, RNA expression, RNA synthesis, antisense; enzymatic nucleic acid, hammerthead ribozyme; MAXzyme; inozyme; inozyme; amberzyme; G-cleaver ribozyme; decoy molecule; aptemer; HBV reverse transcriptase; Enhancer I region; viral replication; degenerative; disease state; HBV infection; HCV infection; cirrhosis; liver failure; hepatocellular carcinoma; hepatotropic; cytostatic; virucide; antiinflammatory; substrate; ss.
                Gape
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               <u>ب</u>
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Lee
                ..
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Mcswiggen J, Morrissey D, Pavco P,
                1; Indels
58.8%; Pred. No. 1.7e+02; ive 6; Mismatches 1
                                                                                                                                                                                                                                             HBV inozyme substrate sequence #118.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Example 1; Page 152; 387pp; English.
                                              544
                                                                         1 ucuućućcakcuaucaak 17
                                                                                                                                                     ACD51886 standard; RNA; 17 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           26-MAR-2001; 2001US-00817879.
08-JUN-2001; 2001US-0087478.
08-JUN-2001; 2001US-0296876P.
24-OCT-2001; 2001US-0335659P.
05-DEC-2001; 2001US-0337055P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                26-MAR-2002; 2002WO-US009187.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RIBOZYME PHARM INC.
BLATT L.
                                            528 TTTTCTGGACTATCAAG
                                                                                                                                                                                                                (first entry)
                10; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Macejak D,
Roberts E;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          BLATT L.
MACEJAK D.
MCSWIGGEN J.
MORRISSEY D.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           WPI; 2003-229207/22.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  PAVCO P.
LEE P.
DRAPER K.
ROBERTS E.
                                                                                                                                                                                                                                                                                                                                                                                                                    Hepatitis B virus.
                                                                                                                                                                                                                                                                                                                                                                                                                                                   WO200281494-A1
                                                                                                                                                                                                               24-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 17-0CT-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Blatt L, I
Draper K,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         infection.
                                                                                                                                                                                   ACD51886;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (MORR/)
(PAVC/)
(LEEP/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (RIBO-)
(BLAT/)
(MACE/)
(MCSW/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (DRAP/)
(ROBE/)
                Matches
                                                                                                                                      ACD51886
                                                                                                                                                                     요
                                              à
```

Lee P;

```
..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    The invention relates to an enzymatic nucleic acid molecule that specifically cleaves RNA derived from hepatitis B virus (HBV) and 2.°Opprising one or more binding arms, without requiring the presence of a 2.°OH group within the molecule for activity. The nucleic acids are useful for treating hepatitis B virus infection, hepatitis, acids are combination with other therapies such liver failure, either alone or in combination with other therapies such as lamivudine and interferons. The nucleic acids are useful as diagnostic tools to examine genetic drift and
that modulate the expression and/or replication of HCV. The compounds and methods of the invention are useful for the treatment of degenerative and disease states related to HBV and HCV infection, replication and gene expression such as circhnosis, liver failure, and hepatocellular carcinoma. The present sequence represents a substrate for one of the HBV ribozyme, inozyme, G-cleaver, zinzyme, DNAzyme or amberzyme sequences
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      specifically cleaving RNA derived from hepatitis B virus and comprising one or more binding arms, useful for treating hepatitis and cirrhosis.
                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Novel enzymatic nucleic acid molecule such as DNAzymes and inozymes
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Hepatitis B virus; HBV; ss; enzymatic nucleic acid; RNA cleavage; hepatitis B virus infection; hepatitis; hepatocellular carcinoma; cirrhosis, liver failure; lamivudine; interferon; genetic drift; virucide; hepatotropic; antiinflammatory; cytostatic.
                                                                                                                                                                                                                                                                                                                                                                                   ..
                                                                                                                                                                                                                                                                                                               0.4%; Score 15.4; DB 1; Length 17; 8.8%; Pred. No. 1.7e+02;
                                                                                                                                                                                                                                                                                                                                                                            1; Indels
                                                                                                                                                                                                                                                        Seguence 17 BP; 4 A; 3 C; 4 G; 0 T; 6 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Hepatitis B virus (HBV) RNA target sequence #819.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ñ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Morrissey
                                                                                                                                                                                                                                                                                                                                                                                6; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Disclosure; SEQ ID NO 819; 122pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Mcswiggen JA,
                                                                                                                                                                                        disclosed in the present invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                530 TICIGGACTATCAAGIT 546
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    92US-00882712.
94US-00193627.
99US-00436430.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20-MAR-2000; 2000US-00531025.
09-AUG-2000; 2000US-00636385.
24-OCT-2000; 2000US-00696347.
08-JUN-2001; 2001US-00877478.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          15-JAN-2003; 2003US-00342902.
                                                                                                                                                                                                                                                                                                                                               58.88;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                :: | | | | | : | : | | | : | : | | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : | : |
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADM58685 standard; RNA; 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (first entry)
                                                                                                                                                                                                                                                                                                                                                                                   10; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    MCSWIGGEN J A. MORRISSEY D.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Blatt L,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2004-247781/23
                                                                                                                                                                                                                                                                                                                                            Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DRAPER K.
BLATT L.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Hepatitis B virus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      US2004054156-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           14-MAY-1992;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        08-NOV-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     07-FEB-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          03-JUN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                18-MAR-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Draper K,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ADM58685
                                                                                                                                                                                                                                                                                                                     Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (DRAP/)
(BLAT/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (MCSW/)
                                                                                                                                                                                                                                                                                                                                                                                   Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 21
ADM58685
      8
                                                                                                                                                                                                                                                                                                                                                                                                                                                ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    요
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    The invention relates to an enzymatic nucleic acid molecule that specifically cleaves RNA derived from hepatitis B virus (HBV) and comprising one or more binding arms, without requiring the presence of a 2'-OH group within the molecule for activity. The nucleic acids are useful for treating hepatitis B virus infection, hepatitis, hepatocellular carcinoma, cirrhosis and liver failure, either alone or in combination with other therapies such as lamivudine and interferons. The nucleic acids are useful as diagnostic tools to examine genetic drift and mutations within diseased cells, for detecting the presence of HBV RNA in
mutations within diseased cells, for detecting the presence of HBV RNA in a cell, for the study of RNA and for down-regulating gene expression of target genes in bacterial, fungal, viral, plant or mammalian cells. This sequence represents an HBV RNA target sequence, used in the scope of the invention. Note: The sequence data for this patent is also available in electronic format from USPTO at seqdata.uspto.gov/sequence.html.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  virus and comprising
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Novel enzymatic nucleic acid molecule such as DNAzymes and inozymes specifically cleaving RNA derived from hepatitis B virus and comprising one or more binding arms, useful for treating hepatitis and cirrhosis.
                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Hepatitis B virus, HBV, ss; enzymatic nucleic acid, RNA cleavage, hepatitis B virus infection; hepatitis; hepatocellular carcinoma; cirrhosis; liver failure; lamivudine; interferon; genetic drift; virucide; hepatotropic; antiinflammatory; cytostatic.
                                                                                                                                                                                                 ;
0
                                                                                                                                                            0.4%; Score 15.4; DB 1; Length 17; 58.8%; Pred. No. 1.7e+02; ive 6; Mismatches 1; Indel8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Hepatitis B virus (HBV) RNA target sequence #1872.
                                                                                                                            Seguence 17 BP; 4 A; 3 C; 4 G; 0 T; 6 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ä
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Morrissey
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Disclosure; SEQ ID NO 1872; 122pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Mcswiggen JA,
                                                                                                                                                                                                                                    530 TTCTGGACTATCAAGTT 546
                                                                                                                                                                                                                                                         ::|:||||:|:||| : uvcuggacuaucaaggu 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        92US-00882712.
94US-00193627.
99US-00436430.
                                                                                                                                                                                                                                                                                                                                                               ADM59738 standard; RNA; 17 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20 MAR-2000; 2000US-00531025.
09-AUG-2000; 2000US-00636385.
24-OCT-2000; 2000US-00696347.
08-JUN-2001; 2001US-00877478.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        15-JAN-2003; 2003US-00342902
                                                                                                                                                                                                                                                                                                                                                                                                                                    (first entry)
                                                                                                                                                                                                   10; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    BLATT L.
MCSWIGGEN J A.
MORRISSEY D.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WPI; 2004-247781/23.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Blatt L,
                                                                                                                                                                                Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Hepatitis B virus.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DRAPER K.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    US2004054156-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            14-MAY-1992;
                                                                                                                                                                                                                                                                                                                                                                                                                                      03-JUN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             08-NOV-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       18-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Draper K,
                                                                                                                                                                                                                                                                                                                                                                                                  ADM59738;
                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (BLAT/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (MORR/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (DRAP/)
                                                                                                                                                                                                                                                                                                                              212
                                                                                                                                                                                  Best Loca
Matches
                                                                                                                                                                                                                                                                                                                                              ADM59738
   8888888888
                                                                                                                                                                                                                                      ð
```

```
RESULT 214
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AAT61999
       셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             8
                                                                                                                                                                                        ö
a cell, for the study of RNA and for down-regulating gene expression of target genes in bacterial, fungal, viral, plant or mammalian cells. This sequence represents an HBV RNA target sequence, used in the scope of the invention. Note: The sequence data for this patent is also available in electronic format from USPTO at seqdata.uspto.gov/sequence.html.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           gene expression; downregulation; interleukin-5; IL-5; ICAM-1; intercellular adhesion molecule; rel A; tumour necrosis factor; TNF-alpha; respiratory syncytial virus; RSV; bcr-abl; oncogene; translocation; chronic myelogenous leukaemia; CML; cancer; Philadelphia chromosome; inflammation; autoimmune disease; atherosclerosis; myocardial infarction; stroke; reetenosis; transplant rejection; rheumatoid arthritis; psoriasis; myocardial ischaemia; Kawasaki disease; septic shock; HIV; human immunodeficiency virus; acquired immune deficiency syndrome; AIDS;
                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human TNF-alpha hairpin ribozyme target sequence (nt position 1178).
                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Enzymatic nucleic acid; ribozyme; trans cleavage; inhibition;
                                                                                                                                                    Length 17;
                                                                                                                                                                                      1; Indels
                                                                                                               Sequence 17 BP; 4 A; 4 C; 3 G; 0 T; 6 U; 0 Other;
                                                                                                                                                  Score 15.4; DB 1;
Pred. No. 1.7e+02;
                                                                                                                                                                                        6; Mismatches
                                                                                                                                                                                                                             544
                                                                                                                                                                                                                                              1 UCUUCUGGACUAUCAAG 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  940S-002181934
940S-0022495
940S-0022795
940S-00228041
940S-00291932
940S-00291932
940S-00291932
940S-00291932
940S-00391090
940S-00311749
940S-00311749
940S-00311749
940S-00311749
940S-00311749
940S-00311749
940S-00311749
940S-00317608
                                                                                                                                                    0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   95WO-IB000156
                                                                                                                                                                                                                         528 TTTTCTGGACTATCAAG
                                                                                                                                                                                                                                                                                                                                                    AAT56721 standard; RNA; 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                   (first entry)
                                                                                                                                                                                      10; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                 (revised)
                                                                                                                                                                   Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                18-MAY-1994;
06-JUL-1994;
15-AUG-1994;
16-AUG-1994;
17-AUG-1994;
19-AUG-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                               25-MAR-2003
02-APR-1997
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WO9523225-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 23-FEB-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  16-DEC-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               15-APR-1994,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              02-SEP-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 08-SEP-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                23-SEP-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          07-APR-1994
15-APR-1994
                                                                                                                                                                                                                                                                                                                                                                                           AAT56721;
                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                   AAT 56 721 / AAT 5
   88888888
                                                                                                                                                                                                                                                        g
```

```
Stinchcomb DT, 'Chowrira B, Direnzo A, Draper KG, Dudycz LW;
Grimm S, Karpeisky A, Kisich K, Matulic-Adamic J, Mcswiggen JA;
Modak A, Pavco P, Beigleman L, Sullivan SM, Sweedler D, Thompson JD;
Tracz D, Usman N, Wincott FB, Woolf T;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The present sequence represents a preferred target sequence for an tenzymatic nucleic acid (i.e. a ribozyme) which cleaves TNF-alpha mRNA at the nucleotide base position indicated in the DE line. Regions of the mRNA that do not form secondary folding structures and that contain potential hammerhead and hairpin ribozyme cleavage sites were identified by computer analysis. Ribozymes directed against these mRNA sequences were designed and synthesised with modifications that improve their nuclease resistance. The ribozymes are designed to cleave the target sequences and thereby inhibit TNF-alpha expression, making them potentially useful for treating rheumatorid arthritis, septic shock and other inflammatory disorders including psoriasis, as well as for treatment of AIDS. (Updated on 25-MAR-2003 to correct PI field.)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     detection; nucleic acid; aqueous solution; length analysis; repeat unit; microsatellite marker; repetitive element; microorganism; probe; genotyping; clinical diagnosis; quantification; sequencing; mutation; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                   Ribozymes having modified bases and methods for producing them - for use in inhibiting disease related genes.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Nucleic acid assay - involving interlinking circular nucleic acid
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ..
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match 0.4%; Score 15.4; DB 1; Length 18; Best Local Similarity 94.1%; Pred. No. 1.9e+02; Matches 16; Conservative 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 18 BP; 3 A; 5 C; 3 G; 0 T; 7 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Probe 3 for Auxis thazard detection.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Claim 2; Page 259; 407pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            3416 TCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ВР
95US-00380734.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       95DE-01033354.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ~
                                                                       (RIBO-) RIBOZYME PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       18 TCAAGGAAGTCTGGAAA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AAT61999 standard; DNA; 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (first entry)
                                                                                                                                                                                                                                                                                                                                       WPI; 1995-351090/45.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 1997-166755/16.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DE19533354-A1.
   30-JAN-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (OZKA/) OZKAN
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            08-SEP-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       08-SEP-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       08-DEC-1997
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           molecules
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAT61999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Ozkan D;
```

ö

10001863-3.81.rng

```
A new method for detection of nucleic acids in aqueous solution is claimed, which may be used for length analysis of repeat unit containing sections of microsatellite markers or other repetitive elements. The method may also be used for microorganism detection, parallel detection of several target nucleic acids using several primer pairs for enzymatic amplification and/or several differently labelled detection probes and/or ard probes, genotype determination, determining relatedness of organisms, clinical diagnosis, food chemistry, anthropology, forensic analysis, specific quantification of nucleic acids, determining the RNA concentration of tumour markers, determining nucleic acid sequences for antigenic determinants, nucleic acid sequence analysis, antigenic determinants genetic changes involved in the development of drug resistance. AATG1997-G2000 are species specific probes for detection of Auxis thazard using the new method
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Oligonucleotide sequences which bind nuclear proteins and surfactants found in lung cells, useful for detecting cancers that originate in the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human; surfactant protein B; SPB; Thyroid transcription factor; TTF-1;
lung cancer; thyroid cancer; 5'SPB-f1; ss; probe; HNF-3; EMSA;
electrophoretic mobility shift assay.
                                                                                                                                                                                                                                                                                                                             ;
0
                                                                                                                                                                                                                                                                                              Length 18;
                                                                                                                                                                                                                                                                                                                             1; Indels
                                                                                                                                                                                                                                                                Sequence 18 BP; 3 A; 4 C; 5 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                              Score 15.4; DB 1;
Pred. No. 1.9e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Human surfactant protein B, SPB, probe 5'SPB-f1.
                                                                                                                                                                                                                                                                                                                              0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Example 2; Fig 10a; 76pp; English.
Example 3; Page 7; 10pp; German.
                                                                                                                                                                                                                                                                                                                                                            3125 TGATGACTGCAGTCGTC 3141
                                                                                                                                                                                                                                                                                                                                                                            뮵.
                                                                                                                                                                                                                                                                                               0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         99US-00320337
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      94US-00245356
95US-00442809
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAS11810 standard; DNA; 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Bohinski RJ, Whitsett JA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (first entry)
                                                                                                                                                                                                                                                                                                              Local Similarity 94.1
es 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (BOHI/) BOHINSKI R J. (WHIT/) WHITSETT J A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WPI; 2001-513959/56.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Oligonucleotide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              US2001016352-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         26-MAY-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        18-MAY-1994;
17-MAY-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    24-OCT-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           23-AUG-2001.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAS11810;
                                                                                                                                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT 215
AAS11810
                                                                                                                                                                                                                                                                                                                              Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         요
```

The invention relates to an oligonucleotide which includes at least 1 nucleic acid sequence which binds to at least 1 nuclear protein found in lung cells (e.g. the thyroid transcription factor 1, TFF-1, protein). The oligonucleotide can be expressed in lung cells via a vector and can be used to target therapeutic agents to kill lung or thyroid cancer cells. The oligonucleotide can be used to detect or diagnose lung or thyroid cancer. The oligonucleotides may be designed from the sequences of, for

ö

Gaps

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    The invention relates to an oligonuclectide which includes at least 1 nucleic acid sequence which binds to at least 1 nuclear protein found in lung cells (e.g. the thyroid transcription factor 1, TTF-1, protein). The oligonuclectide can be expressed in lung cells via a vector and can be used to target therapeutic agents to kill lung or thyroid cancer cells. The oligonucleotide can be used to detect or diagnose lung or thyroid cancer. The oligonucleotides may be designed from the sequences of, for example, the promoters of lung specific genes such as those encoding surfactant proteins. The present sequence is a Human surfactant protein B, SPB, probe 5/fl based on the SPB-fl probe and is used to identify TTF-1 and HNF-3 binding sites in the SPB promoter using EMSA, electrophoretic
                                                                                                                                                                             ö
example, the promoters of lung-specific genes such as those encoding surfactant proteins. The present sequence is a Human surfactant protein B, SPB, probe 5'SPB-fl and is used to identify TTF-1 and HNF-3 binding sites in the SPB promoter using EMSA, electrophoretic mobility shift
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Oligonucleotide sequences which bind nuclear proteins and surfactants found in lung cells, useful for detecting cancers that originate in the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Human; surfactant protein B; SPB; Thyroid transcription factor; TTF-1; lung cancer; thyroid cancer; S'f1; ds; probe; HNF-3; EMSA; electrophoretic mobility shift assay.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ;
                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match 0.4%; Score 15.4; DB 1; Length 18; Best Local Similarity 94.1%; Pred. No. 1.9e+02; Matches 16; Conservative 0; Mismatches 1; Indels
                                                                                                                                         0.4%; Score 15.4; DB 1; Length 18;
4.1%; Pred. No. 1.9e+02;
ve 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 18 BP; 5 A; 5 C; 6 G; 2 T; 0 U; 0 Other;
                                                                                                          Sequence 18 BP; 2 A; 6 C; 5 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                            Human surfactant protein B, SPB, probe 5'fl.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Example 2; Fig 9a; 76pp; English.
                                                                                                                                                                                                                  1798 CCCTCCAGGTTCTTGAT 1814
                                                                                                                                                                                                                                                                                                                                         BP.
                                                                                                                                                                                                                                                   2 cccrccaggracrraar 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            94US-00245356.
95US-00442809.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             99US-00320337.
                                                                                                                                                               94.18;
                                                                                                                                                                                                                                                                                                                                         AAS11808 standard; DNA; 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    JA;
                                                                                                                                                                                                                                                                                                                                                                                                               (first entry)
                                                                                                                                         Ouery Match
Best Local Similarity 94.1'
Matches 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Bohinski RJ, Whitsett
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (BOHI/) BOHINSKI R J. (WHIT/) WHITSETT J A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         mobility shift assay
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2001-513959/56.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         US2001016352-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               26-MAY-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               18-MAY-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              17-MAY-1995;
                                                                                                                                                                                                                                                                                                                                                                                                               24-OCT-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            23-AUG-2001
                                                                                                                                                                                                                                                                                                                                                                            AAS11808;
                                                                                                                                                                                                                                                                                                        RESULT 216
AAS11808/c
                                                                              assay
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              lung.
                                                                                                                                                                                                                                                                                                                                                             8
                                                                                                                                                                                                                                                   임
```

RESULT 217

ઠ g

```
New nucleic acid, useful in imparting disease resistance to a plant or in preparing a composition for treating cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  py. tomato D23000 Avr/Hop proteins, and the polynucleotide sequences encoding them. Also disclosed are expression vectors, host cells, and transgenic plants comprising polynucleotide sequences of the invention. The polynucleotide and polypeptide equences are useful in imparting disease resistance to a plant or in preparing a composition for treating cancer. The sequences may also be used to make a plant hypersusceptible to colonisation by nonpathogenic bacteria, modify a metabolic pathway in a cell, cause eukaryotic cell death, and inhibit programmed cell death. The present sequence represents a PCR primer used in the examples of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             present invention relates to the isolation of Pseudomonas syringae
                                                                                         Avr; Hop; transgenic plant; disease resistance; cancer; bacteria; metabolic pathway; eukaryotic cell death; programmed cell death; cytostatic; PCR; primer; ss.
                                                          Pseudomonas syringae pv. tomato DC3000 Hop gene PCR primer #45.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0.4%; Score 15.4; DB 1; Length 18;
4.1%; Pred. No. 1.9e+02;
ve 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      cytostatic; gene therapy; Avr; Hop; cancer; primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Cartinhour SW, Schneider DJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 18 BP; 5 A; 7 C; 2 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Pseudomonas syringae anti-cancer gene primer #65.
                                                                                                                                                                              Pseudomonas syringae; pv. tomato str. DC3000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Example, SEQ ID NO 192, 209pp, English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 2896 GCATTTCAACCACTCA 2912
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADL12254 standard; DNA; 18 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2 GCATTTCAACCAGCTCA 18
                                                                                                                                                                                                                                                                                                12-FEB-2003; 2003US-00365742.
                                                                                                                                                                                                                                                                                                                                     12-FEB-2002; 2002US-0356408P.
10-MAY-2002; 2002US-0380185P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       94.18;
                  (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                (ALFA/) ALFANO J R.
(CART/) CARTINHOUR S W.
(SCHN/) SCHNEIDER D J.
(TANG/) TANG X.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Collmer A, Alfano JR,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WPI; 2003-875735/81
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
Best Local Similarity
Matches 16; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        present invention.
                                                                                                                                                                                                                     US2003204868-A1.
                                                                                                                                                                                                                                                                                                                                                                                                 COLLMER
                    11-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         06-MAY-2004
                                                                                                                                                                                                                                                          30-OCT-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ADL12254;
                                                                                                                                                                                                                                                                                                                                                                                                 COLL/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     RESULT 219
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ADL12254
ID ADL1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     The present invention relates to nucleic acid probes, which are useful for assaying nucleic acids by hybridising with a target nucleic acid, in which a single-stranded oligonucleotide is labelled with a fluorescent substance and a quencher in a manner that the fluorescence intensity of the hybridisation reaction system is increased after completion of the hybridisation but no stem loop structure is formed. The probes are useful for assaying nucleic acids and their polymorphism and mutation, particularly useful for e.g. analytical applications, disease diagnosis and microbial identification. The present sequence was used to illustrate
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Fluorescently-labeled nucleic acid probes for assaying nucleic acids and their polymorphism and mutation, particularly useful in science and medicine for e.g. analytical applications, disease diagnosis and microbial identification.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   봈
                                                                                                                                                                                                                                                                                                              Probe; polymorphism detection; mutation detection; disease diagnosis; microbial identification; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Yamada
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Kurata S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Score 15.4; DB 1; Length 18;
Pred. No. 1.9e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 18 BP; 14 A; 0 C; 0 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Kamagata Y, Torimura M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (NAAD-) NAT INST ADVANCED IND SCI & TECHNOLOGY.
(XANK-) KANKYO ENG CO LTD.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Example 12; Page 60; 152pp; Japanese.
                                                                                                                                                                                                                                                                             Probe d for assaying nucleic acids
                  CCCTCCAGGTTCTTGAT 1814
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    3321
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        rrrrrrrrrraiai 2
                                                                                                                                                         BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    27-JUN-2000; 2000JP-00193133.
03-AUG-2000; 2000JP-00236115.
26-SEP-2000; 2000JP-00292483.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                27-JUN-2001; 2001WO-IB001147.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      0.4%;
                                                     cccrccaccrccar
                                                                                                                                                       ABL95898 standard; DNA; 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    TTTTTTTTTTTTT
                                                                                                                                                                                                                                   19-JUN-2002 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Kanagawa T,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           WPI; 2002-195876/25.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                  WO200208414-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         the invention
                                                                                                                                                                                                                                                                                                                                                                          Unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                           31-JAN-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             16;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Kurane R,
Yokomaku T;
                1798
                                                     17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    3305
                                                                                                                                                                                              ABL95898;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
```

Tang X;

ö

Gaps

ö

Pseudomonas syringae; pv tomato DC3000.

ADG73198 standard; DNA; 18

RESULT 218

셤 δ

Matches

ADG73198

ADG73198 ID ADG7 XX AC ADG7

WO2003068930-A2.

9.4

×

```
The invention relates to novel Pseudomonas Avr and Hop genes, a sequence that hybridizes with these sequences under stringency conditions comprising a hybridization medium that includes 0.9 x saline sodium citrate (SSC) buffer at a temperature of 42 deg C. The nucleic acid molecule is useful for preparing a composition for treating cancer. This sequence corresponds to a PCR to isolate and amplify one of the genes of
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Measurement of nucleic acids, using a nucleic acid probe and analysis of
                                                                                                                                                                                                        nucleic acid molecule, useful for preparing a composition for
                                                                                                                                                                                                                                                                                                                                                                                           Length 18;
                                                                                                                                                                                                                                                                                                                                                                                                                   1; Indels
                                                                                                                                                          Schneider DJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ss; fluorochrome; nucleic acid probe; fluorescence.
                                                                                                                                                                                                                                                                                                                                                                    Sequence 18 BP; 5 A; 7 C; 2 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (KANK-) KANKYO ENG KK.
(KEIZ-) KEIZAI SANGYOSHO SANGYO GIJUTSU SOGO KEN
                                                                                                                                                                                                                                                                                                                                                                                         Score 15.4; DB 1;
Pred. No. 1.9e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                   Mismatches
                                                                                                                                                                                                                                             Disclosure; SEQ ID NO 192; 284pp; English.
                                                                                                                                                         Cartinhour SW,
                                                                                                       US SEC OF AGRIC.
UNIV NEBRASKA.
UNIV KANSAS STATE RES FOUND.
                                                                                                                                                                                                                                                                                                                                                                                                                   0
                                                                                                                                                                                                                                                                                                                                                                                                                                           2896 GCATTTCAACCACTCA 2912
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20-APR-1999; 99JP-00111601.
24-AUG-1999; 99JP-0023666.
30-AUG-1999; 99JP-00242693.
01-FEB-2000; 2000JP-00028896.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (BIOI-) BIOINDUSTRY KYOKAI SH
                                                                                             (CORR ) CORNELL RES FOUND INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20-APR-2000; 2000JP-00120097.
                                  12-FEB-2003; 2003WO-US004450.
                                                                                                                                                                                                                                                                                                                                                                                           0.4%;
                                                        12-FEB-2002; 2002US-0356408P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  GCATTTCAACCAGCTCA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                   16; Conservative
                                                                                                                                                         Collmer A, Alfano JR,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 2002-134193/18
                                                                                                                                                                                 WPI; 2003-679632/64
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ABA97625 standard;
                                                                                                                                                                                                                                                                                                                                                                                                      Local Similarity
                                                                                                                                                                                                                       treating cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              JP2001286300-A.
                                                                                                                                                                                                                                                                                                                                             the invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Unidentified
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              11-APR-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     16-0CT-2001
          21-AUG-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  N
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ABA97625;
                                                                                                                                                                                                                                                                                                                                                                                            Query Match
                                                                                                        (USDA ) (UYNE-) (UYNE-) (UNIV )
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Probe d.
                                                                                                                                                                                                                                                                                                                                                                                                                   Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                New
요
```

```
The sequences given in AAQ41826-61 represent sequences which are bound in an electrophoretic mobility shift assay (EMSA) by Myc. The isolated consistences contain the central E box core of CAGGTG which binds very weakly with Myc homo-oligomers (C1 complex), but more tightly with Myc homo-oligomers (C2 complex), but more tightly with Myc complex. The C2 complex but more tightly with Myc and c1 in addition to Myc. The additional factor copurifies with Myc and c1 resembles Max protein. A second copurifying 40-50 kD factor has been c1 dentified (forming C2 complex). Sites selected by the C2' complex contain the core CACCTG which bears remarkable homology to a myogenin binding site (see AAQ41763). Oligonucleotides containing the E box can be used in the purification of Myc from a mammalian source. See also capadated target sequences may be used in a method to inhibit c-Myc oncoprotein activity. (Updated on 25-MAR-2003 to correct PN field.)
                                                                                                                                                                                                                                             ó,
                                                                This invention relates to a method for measuring nucleic acids using a nucleic acid probe labelled with a fluorochrome. The nucleic acid probe decreases the fluorescence of the fluorochrome when hybridised with a target nucleic acid, the decrease in the fluorescence is measured. The method can be used for measuring a target nucleic acid
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Prodn. of c-Myc protein from mammalian cells - and detection of c Myc inhibitors for use in cancer therapy.
                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   complex;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Myc; c-myc; mammalian; E box; cancer; therapy; C1; C2; C2'; complex; homo-oligomer; hetero-oligomer; myogenin; Max; oncoprotein; primer; probe; electrophoretic mobility shift assay; EMSA; ss.
                                                                                                                                                                                                                                             .
0
                                                                                                                                                                                                     0.4%; Score 15.4; DB 1; Length 19; 94.1%; Pred. No. 2e+02; ive 0; Mismatches 1; Indels
                                                                                                                                                                       Sequence 19 BP; 15 A; 0 C; 0 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  CHO C2 and C2' complex AT rich binding site #4.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Disclosure, Fig 7b; 101pp; English.
                                 Example 5; Page 17; 34pp; Japanese.
                                                                                                                                                                                                                                                                                  3305 TTTTTTTTTTTTT 3321
                                                                                                                                                                                                                                                                                                                                                                                                            BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  91US-00785567.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  92WO-US008603.
                                                                                                                                                                                                                                                                                                                   N
                                                                                                                                                                                                                                                                                                                                                                                                            AAQ41846 standard; DNA; 20
                                                                                                                                                                                                                                                                                                            18 TTTTTTTTTATATAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (GEHO ) GEN HOSPITAL CORP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                               Local Similarity 94.1
les 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Kingston RE, Papoulas
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (revised)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 1993-167291/20
the obtained data.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WO9308701-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    09-OCT-1992;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     30-OCT-1991;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                25-MAR-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 03-SEP-1993
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               13-MAY-1993
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                               AAQ41846;
                                                                                                                                                                                                              Query Match
                                                                                                                                                                                                                                                                                                                                                                         RESULT 221
                                                                                                                                                                                                                                                 Matches
                                                                                                                                                                                                                                                                                                                                                                                           AAQ41846/
 FX8X55555X8
                                                                                                                                                                                                                                                                                      ઠે
                                                                                                                                                                                                                                                                                                                     g
                                                                                                                                                                                                                                                                                                                                                                                                                              ö
```

Gaps

; 0

Seguence 20 BP; 13 A; 3 C; 1 G; 3 T; 0 U; 0 Other;

10001863-3.81.rng

AAA40987 standard; DNA; 20 BP

AAA40987

Antisense oligonucleotide; phosphorothioate; TNFalpha; cytokine; inhibit; tumour necrosis factor alpha; inflammatory bowel disease; diabetes; rheumatorid arthritis; infectious disease; multiple sclerosis; hepatitis; pancreatitis; atopic dermatitis; allograft rejection; autoimmune disease; inflammatory disease; ss.

Human TNFalpha antisense oligonucleotide ISIS# 100606.

(first entry)

16-AUG-2000

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            PCR primers AA201426-206209 were used to amplify open reading frames CORFS of the genome of Chlamydia trachomatis (see AA201425). These ORFS encode polypeptides (see AAX36754-Y37949) which can be used as vaccines against Chlamydia trachomatis. Antisense and ribozyme sequences can also be used to control growth of the microorganism. Chlamydia trachomatis is responsible for a large number of diseases, e.g. eye diseases such as conventional trachoma, nonendemic trachoma, paratrachoma, and inclusion conjunctivitis; genital diseases such as nongonococcal uretritis, perior protections and properties of the invention may be of use in treating these
                                                                                                                                                                                                                                                                                         Vaccine, eye disease, conventional trachoma, nonendemic trachoma, paratrachoma, inclusion conjunctivitis, genital disease, perihepatitis, nongonococcal uretritis, epidymitis, cervicitis, salpingitis, PCR primer, bartholinitis, pneumopathy, venereal lymphogranulomatosis, ss.
                               Gaps
                               ;
0
   Length 20;
                                                                                                                                                                                                                                                              PCR primer used to amplify an ORF of Chlamydia trachomatis.
                              1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Sequence 20 BP; 9 A; 0 C; 10 G; 1 T; 0 U; 0 Other;
Score 15.4; DB 1;
Pred. No. 2.2e+02;
                              0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Genome sequence of Chlamydia trachomatis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Disclosure; Page 1604; 1755pp; English.
                                                            3315
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              97FR-00015041.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            97FR-00016034.
Query Match
Best Local Similarity 94.1%;
Matches 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 98WO-IB001939
                                                           3299 GATATATTTTATTTT
                                                                                       20 GATGTATTTTTTTTT
                                                                                                                                                                   AAZ03406 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                       Chlamydia trachomatis
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 1999-371125/31.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (GEST ) GENSET
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               27-NOV-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              17-DEC-1997;
04-NOV-1998;
                                                                                                                                                                                                                               07-OCT-1999
                                                                                                                                                                                                                                                                                                                                                                                                                    WO9928475-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              28-NOV-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                   10-JUN-1999.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Griffais R;
                                                                                                                                                                                                                                                                                                                                                                       Synthetic.
                                                                                                                                                                                                 AAZ03406;
                                                                                    g
```

Oligonucleotide for treating diseases associated with human tumor necrosis factor-alpha (TNF-alpha) such as, diabetes and rheumatoid arthritis, comprises nuclectide sequence complementary to intron of nucleic acid encoding TNF-alpha.

Butler MM, Shanahan WJ;

Baker BF, Bennett CF, (ISIS-) ISIS PHARM INC.

WPI; 2000-303808/26.

99WO-US023205, 98US-00166186. 99US-00313932.

05-OCT-1999; 05-OCT-1998; 18-MAY-1999;

13-APR-2000.

WO200020645-A1.

Synthetic.

```
This sequence represents an antisense oligonucleotide sequence which targets a region of the human tumour necrosis factor alpha (INFalpha)

targets a region of the human tumour necrosis factor alpha (INFalpha)

mucleotide sequence. TNFalpha is an important cytokine that plays a role in host defence. It is produced mainly in macrophages and monocytes in response to infection, invasion, injury or inflammation. Overexpression of TNFalpha can result in disease states, particularly in infectious, inflammatory and autoimmune diseases. The invention relates to antisense oligonucleotides, such as that repersented by the present sequence which are capable of modulating the TNFalpha gene expression. The oligonucleotides of prionally have a phosphorothicate backbone, and may also optionally contain at least one 2'-0-methoxyethyl modification. The oligonucleotides are useful for modulating the expression of human TNFalpha in cells and tissues, reducing a human cell inflammatory cresponse, reducing the blood glucose level in a human and treating thuman having a disease or condition associated with TNFalpha include diabetes, inflammatory bowel
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          disease, multiple sclerosis, pancreatitis, rheumatoid arthritis, infectious disease, hepatitis, atopic dermatitis or allograft rejection. The antisense oligonucleotides are also useful for modulating the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The antisense oligonucleotides are also useful for modulating t
function of a selected nucleic acid sequence in adipose tissue
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.4%; Score 15.4; DB 1; Length 20; 94.1%; Pred. No. 2.2e+02; ive 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 8 A; 3 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Example 20; Page 95; 283pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                3416 TCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1 TCAAGGAAGTCTGGAAA 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Ouery Match 0.4
Best Local Similarity 94.1
Matches 16; Conservative
ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       셤
```

RESULT 224 AAA10948/c

0; Gaps

Score 15.4; DB 1; Length 20; Pred. No. 2.2e+02; 0; Mismatches 1; Indels

0.4%; 94.18; 2477 GCAGAAGGTGGAGAAGA 2493

ሯ

e

16; Conservative

Best Local Similarity Matches 16; Conserv

Query Match

RESULT 223

AAA10948 standard; DNA; 20

```
2225
                                                                                             (UYRO-) UNIV ROTTERDAM ERASMUS
                                                                                                                                                                                                                                                                           BP
                                                                            99WO-EP007719.
                                                                                     98GB-00019769.
                                                                                                                                                                                                                            ch 0.4%;
1 Similarity 94.1%;
16; Conservative
                                                                                                                                                                                                                                             2209 GCCAGGATGAGGACTGG
                                                                                                                                                                                                                                                      19 GCCAGGATGAGGGCTGG
                                                                                                                                                                                                                                                                           AAD17705 standard; DNA; 20
                                                                                                                                                                                                                                                                                            (first entry)
                (first entry)
                                                                                                              WPI; 2000-271470/23.
                                                                                                                                                                                                                                Local Similarity
                                                           WO200015839-A1
                                                                                     10-SEP-1998;
                                                   Homo sapiens
                                                                            10-SEP-1999;
                14-JUL-2000
                                                                                                                                                                                                                                                                                            10-DEC-2001
                                                                    23-MAR-2000
                                                                                                                                                                                                                                                                                   AAD17705
        AAA10948
                                                                                                                                                                                                                            Query Match
                                                                                                                                                                                                                                    Matches
                                                                                                                                                                                                                                                                  RESULT 225
                                                                                                                                                                                                                                                                       à
                                                                                                                                                                                                                                                    g
```

detection; public hygiene; food evaluation; food poisoning evaluation; PCR primer; ss. Oligonucleotide corresponding to human nucleic acids. Sequence 20 BP; 4 A; 4 C; 3 G; 9 T; 0 U; 0 Other; Saitoh J; Disclosure; Page 31; 50pp; English 1089 AATGTTTCTTCATTTTC 1105 ABQ78421 standard; DNA; 20 BP 97US-00949344. 16-MAR-2001; 2001EP-00106364. 17-MAR-2000; 2000JP-00081806. 31-MAY-2000; 2000JP-00166503. 31-MAY-2000; 2000JP-00166504. 31-MAY-2000; 2000JP-00166505. 0.4%; 94.1%; 97US-00949344. 2000JP-00081805 4 AATGATICITCATITIC (first entry) Ishizuka T, Ishiguro T, 16; Conservative Vibrio parahaemolyticus parahaemolyticus trhl WPI; 2001-572721/65. Query Match Best Local Similarity Matches 16; Conserv (TOYJ ) TOSOH CORP. Homo sapiens US6406846-B1 14-OCT-1997; EP1134292-A2 17-MAR-2000; 05-NOV-2002 14-OCT-1997; 19-SEP-2001 AB078421; RESULT 226 ABQ78421/c ઠ 셤 This sequence represents a PCR primer used to amplify intron 1 of the collagen I-alpha-1 gene. The PCR product is used in the method of the invention for determining susceptibility to bone damage. The method comprises screening for polymorphisms in the vitamin D receptor (VDR) gene or the Collagen I-alpha-1 gene. The the presence of alleles A and/or I, and especially the haplotype baT, are associated with an increased risk of bone damage and a higher risk of bone fracture. The methods are used for determining the susceptibility to bone damage and osteoporosis. Done damage may be any form of structural damage, including fractures, breaks or chips. Identification of those at risk using the method of the invention, allows for preventative measures to be taken, such as modifications to lifestyle, regular exercise, and changes in diet to strengthen bones, and hormone therapy. The present invention allows more effective preventative measures to be taken ; 0 PCR primer; vitamin D receptor; susceptibility; bone damage; polymorphism; increased risk; osteoporosis; structural damage; fracture; break; chips; hormone therapy; prevention; collagen I-alpha-1; ss. Novel methods for determining the susceptibility to bone damage by screening for polymorphisms in the vitamin D receptor gene or collagen Ialphal gene. Gaps Thermostable direct haemolysin-related haemolysin gene; trhl; trh2; thermostable direct haemolysin gene; tdh2; clinical examination; ; 0 Length 20; Primer #2 for collagen Ialphal gene fragment amplification. 1; Indels Sequence 20 BP; 2 A; 10 C; 4 G; 4 T; 0 U; 0 Other; Score 15.4; DB 1; Pred. No. 2.2e+02; V. parahaemolyticus trhl amplifying primer #30. Pols HAP; 0; Mismatches Uitterlinden AG, Van Leeuwen JPTM, Example 1; Page 9; 37pp; English.

```
amplification of a gene selected from the group consisting of Vibrio parahaemolyticus direct haemolysin-related thermostable haemolysin genes (trh) and trh2) and V. parahaemolyticus thermostable direct haemolysin gene (tdh2) or RNA derived therefrom. These oligonucleotides are useful as probe and primer for detection or amplification of V. parahaemolyticus examination, public hygiene, food evaluation or food poisoning evaluation. The present sequence is a primer used to amplify V.
New oligonucleotides as primers and probes, useful for detection or amplification of Vibrio parahaemolyticus thermostable direct hemolysin-related hemolysin genes or RNA derived from them.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human; hereditary pancreatitis; HP; mutation; trypsinogen gene; ss.
                                                                                                                                                                                                                                                                            invention relates to oligonucleotides used for detection or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               .;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Score 15.4; DB 1; Length 20;
Pred. No. 2.2e+02;
0; Mismatches 1; Indels
```

.; 0

```
Determining susceptibility of human patient to hereditary pancreatitis comprises analyzing nucleic acid from patient for presence of mutation in third exon of cationic trypsinogen gene that indicates hereditary pancreatitis.
                                                                                                                                                                                                        The specification describes a method for determining whether a human patient is susceptible to hereditary pancreatitis (HP). The method comprises obtaining nucleic acid from the human patient and analysing the nucleic acid to identify the presence of a single G to A transition mutation in codon 117 in a third exon of a cationic trypsinogen gene, or a single A to T transition mutation at codon 21 in second exon of a cationic trypsinogen gene, that indicates HP. The method is useful for determining whether a human patient is susceptible to HP. The present sequence represents an oligonucleotide, which is used in the course of
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Tumour necrosis factor alpha; TNF-alpha; antiinflammatory; antirheumatic; antidiabetic; dermatological; hepatotropic; antidiabetic; inflammatory bowel disease; Crohn's disease; colitis; rheumatory at all disease; colitis; rheumatoid arthritidabetic; alphacetics; pancreatitis; multiple sclerosis; atopic dermatitis; aschma; hepatitis;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Tumour necrosis factor alpha antisense oligonucleotide #218.
                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15.4; DB 1; Length 20; 94.1%; Pred. No. 2.2e+02; tive 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                            Sequence 20 BP; 7 A; 6 C; 5 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Shanahan WR;
                                Gorry MC;
                                                                                                                                                                         Disclosure; Col 115; 66pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Butler MM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1098 TCATTTTCCTGGTGAG 1114
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         02-APR-2001; 2001US-00824322.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        98US-00166186.
99US-00313932.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               TCATTTCCCTGGTGGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACD05215 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          05-AUG-2003 (first entry)
                                .
B
                                                                                                                                                                                                                                                                                                                                                                                                                                           Best Local Similarity 94.1
Matches 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        antisense technology; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Baker BF, Bennett CF,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (BAKE/) BAKER B F.
(BENN/) BENNETT C F.
(BUTL/) BUTLER M M.
(SHAN/) SHANAHAN W R.
                                Ehrlich
                                                             WPI; 2002-581937/62
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2003-447433/42.
(WHIT/) WHITCOMB D.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         US2003022848-A1.
                                                                                                                                                                                                                                                                                                                                                               the invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        05-OCT-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         18-MAY-1999;
                                ď
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         30-JAN-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           19
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACD05215;
                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 227
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          SXCCCCCCCCXXXTTTTXXXXXIX
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ઠ
```

Treating inflammatory disorders such as inflammatory bowel disease,

ö

Gaps

ö

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Determining susceptibility to bone damage by determining polymorphisms in the methyltetrahydrofolate reductase and collagen I alpha 1 genes or serum levels of homocysteine allows preventing or reducing risk by folic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The invention relates to new methods for determining susceptibility of a subject to bone damage comprise determining which allele(s) of polymorphisms in methyltetrahydrofolate reductase (WTHFR) and collagen lalphal are present in the subject, or measuring level of serum homocysteine in the subject, where elevated level compared to a reference population indicates susceptibility. Also included are determining the susceptibility to bone damage in a subject identified as having the s
                                                                                                          The invention describes a method of treating an inflammatory disorder in an individual, comprising administering to the individual an oligonaclectide upto 30 nuclectides in length complementary to a nucleic acid molecule encoding human tumor necrosis factor (TNF)-alpha. The method is useful for treating an inflammatory disorder such as inflammatory bowel disease, Crohn's disease, colitis or rheumatoid arthritis, in an individual. The method is also useful for treating diabetes, pancreatitis, multiple sclerosis, atopic dermatitis, asthma, and happatitis in an individual. This sequence represents an antisense oligonucleotide used to modulate expression of tumour necrosis factor
Crohn's disease or rheumatoid arthritis, in a subject, by administering oligonucleotide which inhibits expression of human tumor necrosis factor alpha.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human, 88; collagen IalphaI, SNP, single nucleotide polymorphism;
osteoporosis; bone damage; serum homocysteine; folic acid; PCR; primer.
                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                             Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                  1; Indels
                                                                                                                                                                                                                                                                                                                                          Sequence 20 BP; 8 A; 3 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                         Score 15.4; DB 1;
Pred. No. 2.2e+02;
0; Mismatches 1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Disclosure; SEQ ID NO 7; 25pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human collagen Ialphal PCR primer #2.
                                                                             Example 22; Page 37; 142pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Van Meurs JBJ;
                                                                                                                                                                                                                                                                                                                                                                                                                                                    3416 TCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1 rchaechaercrechan 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ADF60910 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         11-OCT-2002; 2002US-00270714.
                                                                                                                                                                                                                                                                                                                                                                               0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                11-OCT-2001; 2001US-0328929P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                         Query Match 0.4
Best Local Similarity 94.1
Matches 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (UITT/) UITTERLINDEN A G. (VMEU/) VAN MEURS J B J.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2003-898055/82.
                                                                                                                                                                                                                                                                                                     alpha (TNF-alpha)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Uitterlinden AG,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     US2003165928-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            treatment
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                12-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      04-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADF60910;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 228
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            acid
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ADF60910/
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       유
```

```
allele of the Spl polymorphism in collagen Ialphal, preventing or reducing susceptibility to bone damage in a subject (comprising prescribing or administering folic acid to a subject at risk for bone damage), preventing or reducing bone damage (comprising determining that a subject as increases susceptibility to bone damage, and prescribing or administering folic acid to the subject, and predicting response of subject to treatment (comprising determining which alleles of the MTHFR and/or collagen Ialphal are present). The invention is useful to prevent or reduce risk of bone damage (e.g. osteoporosis) by treating subjects determined to be susceptible with folic acid. The present sequence is a per primer used to amplify the polymorphic region of the human collagen
                                                                                                                                                                                                                                                                                                                                                   IalphaI gene.
      8
```

Sequence 20 BP; 2 A; 10 C; 4 G; 4 T; 0 U; 0 Other;

```
.;
0
 Length 20;
                    1; Indels
Score 15.4; DB 1;
Pred. No. 2.2e+02;
0; Mismatches 1;
                                         2225
  0.4%;
                                         GCCAGGATGAGGACTGG
                     16; Conservative
           Local Similarity
                                         2209
  Query Match
                     Matches
                                          ð
```

ô

Gaps

sccassarsassecres 3 13 유

Human oligonucleotide sequence. ABZ87410 standard; DNA; 20 17-OCT-2003 (first entry) ABZ87410; RESULT 229 ABZ87410, 

BP

Human; antisense; lung dysfunction; nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds.

Homo sapiens

WO200285308-A2.

31-OCT-2002.

23-APR-2002; 2002WO-US013135.

24-APR-2001; 2001US-0286137P.

(EPIG-) EPIGENESIS PHARM INC

Katz E, Pabalan J, Aguilar D; Li Y, Sandrasagra A, Ka Tang L, Shahabuddin S; Nyce JW, I Miller S,

WPI; 2003-229219/22.

Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or ubiquinone

Disclosure; SEQ ID NO 2652; 872pp; English.

The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation codon, coding region, 5' or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiasthmatic, hypotensive,

immunosuppressive, and cytostatic activity. The composition may have a use in antisense gene therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of an antiinflammatory steroid in a subject, for reducing or depleting levels of, or reducing sensitivity to adenosine, reducing levels of adenosine receptor, producing bronchoditation, increasing levels of ubiquinone or lung surfactant in a subject's tissue, or treating bronchoconstriction, lung allergies, or a respiratory disease or condition. Note: The sequence data for this patent is not represented in the printed specification, but was obtained in electronic format directly from WIPO at ftp.wipo.int/pub/published\_pct\_sequences 

Sequence 20 BP; 6 A; 2 C; 3 G; 9 T; 0 U; 0 Other;

Gaps . 0.4%; Score 15.4; DB 1; Length 20; 4.1%; Pred. No. 2.2e+02; ve 0; Mismatches 1; Indels Query Match
Best Local Similarity 94.1%;
Matches 16; Conservative

ö

ABD23640 standard; DNA; 20 RESULT 230 ABD23640/

ВЪ. 29-JUL-2004 (first entry) ABD23640; 

Human myosin X-derived oligonucleotide SEQ ID 2652.

Human; antisense; bronchoconstriction; allergy; hyposecretion; pain; respiratory tract inflammation; adenosine sensitivity; lung; cancer; surfactant depletion; antialergic; antiinflammatory; antiasthmatic; analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory districtses syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer.

Homo sapiens.

WO200285309-A2.

31-OCT-2002.

23-APR-2002; 2002WO-US013143.

24-APR-2001; 2001US-0286036P.

(EPIG-) EPIGENESIS PHARM INC.

Katz E, Pabalan J, Aguilar D; Li Y, Sandrasagra A, Ka Tang L, Shahabuddin S; Miller S, Nyce JW,

WPI; 2003-093058/08.

ţ Pharmaceutical composition for treating asthma, has antisense oligonuclectide containing less percentage of adenosine, targeted trucleic acids associated with lung airway or lung dysfunction, and bronchodilating agent.

Claim 15; SEQ ID NO 2652; 763pp; English.

This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating bronchoconstriction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene encoding or regulating

```
The invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonucleotides, (c) device, in separate containers, (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antiinflammatory, antiasthmatic, analgesic, hypotensive, immunosuppressive and cytostatic activity, is a composition camprises oligo and is administered to reduce the production comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypeptide present in the lungs. The pulmonary obstruction, and/or bronchoconstriction and/or lung with a disease or condition such as pulmonary vasoconstriction, and or surfactant hypoproduction are associated with a disease or condition such as pulmonary vasoconstriction, allergies and/or surfactant hypoproduction, respiratory distress syndrome, pain, cystic fibrosis, allergic rhinitis, pulmonary transplantation rejection, pulmonary infections, bronchists or cancer.

The reduced adenosine content of the anti-sense oligos corresponding to the oligonucleotides into products that free adenosine into the system e.g., lung, brain, heart, kidney, etc, tissue environment and thereby, to prevent any unwanted effects due to it
   expression of a target polypeptide associated with lung airway or lung
                               dysfunction or cancer and can be anti-sense to the corresponding mRNA
```

Sequence 20 BP; 6 A; 2 C; 3 G; 9 T; 0 U; 0 Other;

Gaps ö Length 20; 1; Indels Score 15.4; DB 1; Pred. No. 2.2e+02; 0; Mismatches 1; 0.4%; 94.1%; 16; Conservative Best Local Similarity Matches 16; Conserv Query Match

ö

1425 TTAGAACAACTAGAACA 1441 TTAGAACAACTAGAATA 1 11 g ઠ

ADH65191 standard; DNA; 20 RESULT 231 ADH65191 

ADH65191;

25-MAR-2004 (first entry)

Human glucocorticoid receptor-specific antisense oligonucleotide #2025.

antisense oligonucleotide; glucocorticoid receptor; infection; inflammation; tumour formation; diabetes; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss;

phosphorothioate backbone; 2'-methoxyethyl; 2'-MOE.

Homo sapiens

WO2003099215-A2.

04-DEC-2003.

20-MAY-2003; 2003WO-US016084

20-MAY-2002; 2002US-0381857P.

(PHAA ) PHARMACIA CORP

Crosby SD, Nalseth AE,

WPI; 2004-035034/03.

New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.

Claim 4; SEQ ID NO 2025; 985pp; English

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
The invention comprises an antisense oligonucleotides that are targeted to nucleic acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  The invention comprises an antisense oligonucleotides that are targeted antisense acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  antisense oligonucleotide; glucocorticoid receptor; infection; infidammation; tumnour formation, diabetes; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2'-methoxyethyl; 2'-MOE.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Human glucocorticoid receptor-specific antisense oligonucleotide #4395.
                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0; Gaps
                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                 0.4%; Score 15.4; DB 1; Length 20;
4.1%; Pred. No. 2.2e+02;
ve 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.4%; Score 15.4; DB 1; Length 20; 94.1%; Pred. No. 2.2e+02; tive 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Sequence 20 BP; 13 A; 1 C; 0 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                             Sequence 20 BP; 3 A; 6 C; 3 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Claim 4; SEQ ID NO 4395; 985pp; English.
                                                                                                                                                                                                                                                                                                                                      1323 TICAAAGGIIGCIGIIC 1339
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   561/c
ADH67561 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                               20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20-MAY-2002; 2002US-0381857P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20-MAY-2003; 2003WO-US016084.
                                                                                                                                                                                                                                         Query Match
Best Local Similarity 94.1%;
Marches 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                               4 rrcaaargrigcrgrrc
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
Best Local Similarity 94.1
Matches 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Crosby SD, Nalseth AE;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (PHAA ) PHARMACIA CORP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2004-035034/03
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      25-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              04-DEC-2003.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADH67561;
                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 232
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ADH67561/
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          888888888888888
                                                                                                                                                                                                                                                                                                                                      ò
```

```
numan; single nucleotide polymorphism; SNP; ss; primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         US2004002151-A1
                                                   JP2003259875-A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Key
modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            17-JUN-2004
                                                                        16-SEP-2003.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Synthetic.
                              Synthetic
                                                                                                                                                                                                                                                                                                                                                                                              N
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ADN06176;
                                                                                                                                                                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 235
                                                                                                                                                                                                                                                                                                                                                     Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                        ADN06176
                                                                                                                                                                                                                                                                                                                                                                                                                                                              ઠે
                                                                                                                                                                                                                                                                                                                                                                                            g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                          The invention comprises an antisense oligonucleotides that are targeted to nucleic acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NoTE: The present sequence contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                             New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
                                                                                                                                                                    antisense oligonucleotide; glucocorticoid receptor; infection; inflammation; tumour formation; diabetes; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2'-methoxyethy1; 2'-MOE.
                                                                                                                                                 Human glucocorticoid receptor-specific antisense oligonucleotide #4452
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15.4; DB 1; Length 20; 94.1%; Pred. No. 2.2e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 12 A; 1 C; 0 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                         Claim 4; SEQ ID NO 4452; 985pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     3300 ATATATTTTTTA 3316
            3300 ATATATTTATTTA 3316
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Primer of the invention #1236
                                                                                    BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ᠬ
                                                                                                                                                                                                                                                                                        20-MAY-2003; 2003WO-US016084.
                                                                                                                                                                                                                                                                                                             20-MAY-2002; 2002US-0381857P.
                               ATATATTTTTATA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADK95516 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADH67618 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (first entry)
                                                                                                                              25-MAR-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 16; Conservative
                                                                                                                                                                                                                                                                                                                                                        Nalseth AE;
                                                                                                                                                                                                                                                                                                                                   (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match
Best Local Similarity
Matches 16; Conserval
                                                                                                                                                                                                                                                                                                                                                                            WPI; 2004-035034/03.
                                                                                                                                                                                                                                                WO2003099215-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       06-MAY-2004
                                                                                                                                                                                                                             Homo sapiens
                                                                                                                                                                                                                                                                    04-DEC-2003.
                                                                                                                                                                                                                                                                                                                                                        Crosby SD,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         13
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADK95516;
                                 20
                                                                                                          ADH67618;
                                                               ADK9551
ID AD
XX AD
AC AD
XX
DT 06
XX DE PE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         셤
                                d
            ò
```

```
1...20
/t-tag= b
/mod_base= OTHER
/note= "Phosphorothioate backbone in which all cytidines
                                                                                                                                                                                                                                                                                                                                                        The present invention relates to a polynucleotide isolated from a human gene and is useful for detecting a single nucleotide polymorphism in a human gene or for diagnosing of disease. The invention enables the detection of a single nucleotide polymorphism in a human gene. The present sequence represents a primer of the invention.
                                                                                                                                                                                                         DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Selenophosphate synthetase 2; SPS2; rheumatoid arthritis; infection; inflammation; tumour; antisense therapy; human; antisense; phosphorothioate backbone; ss.
                                                                                                                                                                                                   Novel polynucleotide useful for PCR amplification along with two lingment from another set of sequences, or for detecting single nucleotide polymorphism in human gene.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      0.4%; Score 15.4; DB 1; Length 20; 94.1%; Pred. No. 2.2e+02; ive 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human SPS2 specific antisense oligonucleotide, ISIS 138245
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /*tag= c
/mod_base= OTHER
/note= "2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          /mod_base= OTHER
/note= "2'-methoxyethyl nucleotides"
16. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 20 BP; 1 A; 7 C; 5 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                              Claim 2; SEQ ID NO 4545; 2627pp; Japanese.
                                                                                                     (KAGA-) KAGAKU GIJUTSU SHINKO JIGYODAN
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        are 5-methylcytidines"
1. .5
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2117
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   08-MAR-2002; 2002JP-00064373.
                                                    08-MAR-2002; 2002JP-00064373.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2101 TIGTAGCAGTICTGGTC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADNO6176 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         /*tag= a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Conservative
                                                                                                                                                       WPI; 2004-093977/10.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Local Similarity
nes 16; Conserv
```

ó;

```
The invention relates to antisense compounds, compositions and methods for modulating the expression of selenophosphate synthetase 2 (SPS2). The composition comprises antisense oligonuclectides targeted to SPS2 gene. The antisense oligonuclectide is useful for modulating the expression of SPS2 in cells or tissues to treat diseases associated with their expression, e.g. rheumatoid arthritis, infections, inflammation or tumours. It is also used for diagnostics, prophylaxis, or as research reagents or kits. The antisense oligonuclectide is useful in antisense therapy. The present sequence is an antisense oligonuclectide targeted to human SPS2 DNA. This sequence is used in the exemplification of the
                                                                                                                                                                                              New antisense oligonucleotides for modulating selenophosphate synthetase 2 (SPS2) expression, useful for diagnosing, preventing or treating conditions associated with SPS2, e.g. rheumatoid arthritis, inflammation
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             cytostatic, cancer, antisense, human telomerase, cell proliferation,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human telomerase antisense sequence related oligonucleotide #5
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 3 A; 8 C; 4 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                      Example 15; SEQ ID NO 20; 47pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               194 GGAGCCTCAGCCCTTCA 210
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             GCAGCCTCAGCCCTTCA 18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ADP88421 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             06-DEC-2002; 2002DE-01058117.
07-FEB-2003; 2003DE-01006084.
                                           28-JUN-2002; 2002US-00186157
                                                                         28-JUN-2002; 2002US-00186157
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 08-DEC-2003; 2003WO-DE004114
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  09-SEP-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (UYDR ) UNIV DRESDEN TECH
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Schmidt U,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Local Similarity 94.1
les 16; Conservative
                                                                                                       (ISIS-) ISIS PHARM INC
                                                                                                                                      Freier SM
                                                                                                                                                                 WPI; 2004-070740/07
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2004-468865/44.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WO2004053116-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            apoptosis; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Schwenzer B,
               01-JAN-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   24-JUN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                            invention.
                                                                                                                                                                                                                                            or tumors
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ADP88421
                                                                                                                                    Watt AT,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Best Loca
Matches
ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          셤
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       /mod_base= OTHER
/note= "Phosphorothioate backbone and all cytidines are 5
-methylcytidines"
New polynucleotide directed against the gene for telomerase catalytic subunit, useful for diagnosis and treatment of solid tumors and leukemia, interacts with specific regions of the mRNA.
                                                                                                                                                    least two target sequence regions, i.e. 2176-2250 and 2296-2392 of the sequence AF015950. The polynucleotides are used for diagnosis, prophylaxis, treatment, monitoring (of progression or therapy) and/or secondary treatment of diseases associated with growth, differentiation and/or division of calls, especially a very wide range of solid cancers and leukaemia, or their metastases, e.g. cancers of the urogenital or gastrointestinal tracts, liver, breast, prostate and bladder. They can also be used to inhibit vitality or proliferation rates of cells, to induce apoptosis and/or cause cell-cycle arrest. The present sequence is an oligonucleotide used in the exemplification of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        death-associated protein kinase-rel. apoptosis-inducing protein kinase; serine/threonine kinase 17B; STK17B; apoptosis; degenerative disorder; neurological disorder; Alzheimer's disease; Parkinson's disease; Amylotrophic lateral sclerosis; ALS; retinitis pigmentoss; blood cell disorder; cancer; autoimmune disorder; viral infection; gene therapy; hyperproliferative disorder; chromosome 2.
                                                                                                   The present invention relates to a polynucleotide, directed against a gene that encodes a catalytic subunit of human telomerase, which interacts with the mRNA of the catalytic subunit of telomerase in at
                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                     0.4%; Score 15.4; DB 1; Length 20;
94.1%; Pred. No. 2.2e+02;
lve 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human DRAK2 antisense oligonucleotide ISIS182454.
                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 5 A; 3 C; 7 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /*tag= c
/mod_base= OTHER
/note= "2'-methoxyethyl residue"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /not\bar{e}= "2'-methoxyethyl residue'
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Location/Qualifiers
                                                                    Example 2; Page 26; 40pp; German.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                              484
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADP68897 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             17 AGCCACGTCTCTACCTT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         12-DEC-2002; 2002US-00318819.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            12-DEC-2002; 2002US-00318819.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Human; 88; antisense; DRAK2;
                                                                                                                                                                                                                                                                                                                                                                                                      94.18;
                                                                                                                                                                                                                                                                                                                                                                                                                                                            468 AGCCACCTCTCTACCTT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              mod_base=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                        Matches 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   .20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                *tag=
                                                                                                                                                                                                                                                                                                                                                                                                       Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      US2004115645-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 nodified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       09-SEP-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        17-JUN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ADP68897;
                                                                                                                                                                                                                                                                                                                                                                                         Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   237
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 23
ADP68897
 ###X#X555555555555
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
```

Gaps

ö

Score 15.4; DB 1; Length 20; Pred. No. 2.2e+02; 0; Mismatches 1; Indels

0.48; 94.1%; Fuessel S, Meye A;

Kraemer K,

Wirth MP,

```
The invention relates to a new compound (e.g. an antisense oligonuclectide), having a sequence comprising 8-80 bp targeted to a nucleic acid encoding DRAKZ (death-associated protein kinase-related apoptosis-inducing protein kinase 2, also known as serine/threonine kinase 17B, STK17B), specifically hybridises with the nucleic acid encoding DRAKZ (appearing as ADF68859 and representing bases 58695-149492 of human chromosome 2) and inhibits expression of DRAKZ. Also included are inhibiting the expression of DRAKZ, in cells or tissues, screening for a modulator of DRAKZ, a diagnostic method for identifying a disease of state, a kit or assay device comprising the compound and treating an animal having a disease or condition associated with DRAKZ. The coligonucleotide compound is useful for preparing a composition for treating hyperproliferative disorders, degenerative disorders, contential statemer's disease, parkinson's disease, controphic lateral sclerosis (ALS), retinitis pigmentosa, blood cell disorders, cancer, autoimmune disorders and viral infection. The present sequence represents an antisense oligonucleotide targeting DRAKZ.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                antimicrobial; antidiabetic; antirheumatic; antiarthritic; gastrointestinal; antiinflammatory; neuroprotective; dermatological; virucide; hepatotropic; human; TNF-alpha; tumour necrosis factor alpha; survivin; TNF-alpha associated disorder; infection; diabetes; rheumatodi archritis; crohn's disease; pancreatitis; multiple sclerosis; atopic dermatitis; hepatitis; antisense oligonucleotide; sclerosis; antisense technology; ss.
                                                                                                                       New oligonucleotide compound that inhibits expression of DRAK2, usefu for preparing a composition for treating hyperproliferative disorder, e.g., cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%; Score 15.4; DB 1; Length 20;
94.1%; Pred. No. 2.2e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Human TNF alpha antisense oligonucleotide seqid 228.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 20 BP; 6 A; 6 C; 3 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0; Mismatches
                                                                                                                                                                                                              Example 15; SEQ ID NO 43; 87pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         2897 CATTICAACCAACTCAG 2913
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               05-OCT-1998; 98US-00166186.
18-MAY-1999; 99US-00313932.
02-APR-2001; 2001US-00824322.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             29-AUG-2003; 2003US-00652795.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              94.1%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ADQ29297 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           CATTTCAGCCAACTCAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         07-OCT-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   16; Conservative
                  (ISIS-) ISIS PHARM INC
                                                        Bennett CF, Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Local Similarity
                                                                                          WPI; 2004-449384/42
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (BAKE/) BAKER B F.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  US2004142346-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         22-JUL-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADQ29297;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 238
ADQ29297
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Matches
8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           g
```

```
The invention describes a double stranded RNA compound 8-80 nucleobases in length targeted to a nucleic acid molecule encoding human TNF-alpha, where the compound specifically hybridises with the nucleic acid molecule encoding TNF-alpha and inhibits the expression of survivin. Also described is a double stranded RNA compound having a fully defined sequence of 20 bp (SEQ ID NO: 432) as given in the specification. Also disclosed are TNF-alpha polypeptides, host cells, vectors and antibodies used in the methods of the invention. The methods and compositions of the present invention are useful for the diagnosis, prevention and/or treatment of diseases or conditions associated with aberrant expression or activity of TNF-alpha, such as infection, diabetes, rheumatoid arthritis, Crohn's disease, pancreatitis, multiple sclerosis, atopic dermatitis and hepatitis. This sequence represents a human tumour necrosis factor alpha (TNF-alpha) antisense oligonucleotide.
                                                                                                                     and survivin, useful for diagnosing, preventing or treating infection, diabetes, arthritis, multiple sclerosis and hepatitis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human; tumour necrosis factor alpha; TNFalpha; ss;
antisense gene therapy; inflammatory disorder; phosphorothioate linkage;
mebylene (methylimino) intersugar linkage; infection; autoimmune disease;
diabbetes; rheumatoid arthritis; Crohn's disease; pancreatitis;
multiple sclerosis; atopic dermatitis; inflammatory bowel disease;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Antisense oligonucleotide targeting human TNFalpha ISIS100606
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15.4; DB 1; Length 20; 94.1%; Pred. No. 2.2e+02; ive 0; Mismatches 1; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         /note= "Phosphorothioate linkages"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 8 A; 3 C; 5 G; 4 T; 0 U; 0 Other,
                                                               Shanahan WR;
                                                                                                                                                                                               Example 22; SEQ ID NO 228; 156pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Location/Qualifiers
                                                               Butler MM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             mod base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ADR02813 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1 rcaaggaagrcrggaaa 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            2003US-00647918
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           98US-00166186,9932,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              3416 TCAAGGAAGTATGGAAA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               đ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Local Similarity 94.1
nes 16; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ...20
*tag=
                                                               Bennett CF,
             (BUTL/) BUTLER M M. (SHAN/) SHANAHAN W R.
BENNETT C F.
                                                                                             WPI; 2004-552557/53.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              colitis; hepatitis
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             US2004152652-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            26-AUG-2003;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           05-OCT-1998;
18-MAY-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             21-OCT-2004
                                                                Baker BF,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ADR02813;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
(BENN/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               RESULT 239
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ADR02813
   ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ô
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ;
0
```

ö

Ë

Summer

```
This sequence represents a primer which was used in the quantitation of the distribution of prostaglandin E2 EP3 III polypeptide. The prostaglandin E2 EP3 III polypeptide and corresponding nucleic acid may be used for screening therapeutic agents for treating disease such as hematological diseases, cardiovascular diseases, usological diseases, cancer, or respiratory diseases in a mammal. The methods comprise detecting binding of a test compound to a prostaglandin E2 EP3 in polypeptide or polymeticaledide, determining activity of prostaglandin E2 EP3 III polypeptide at different concentrations of test compound; or determining activity of prostaglandin E2 EP3 III polypeptide. The regulators of prostaglandin E2 EP3 III polypeptide.

CC presence of a regulator of prostaglandin E2 EP3 III polypeptide. The regulators of prostaglandin E2 EP3 III seases, metabolic diseases, endocrinological disease, metabolic cancer, or respiratory diseases, gastroenterological diseases, and shall for the disease, endocrinological diseases, metabolic cancer, or respiratory diseases, metabolic cancer, or respiratory diseases, and mammal. They are also useful for the disease.
                                                                                                                                                                                                                                                                                                                                          Screening agents for treating, e.g., cancer by detecting binding of compound to prostaglandin B2 EP3 III protein or polynucleotide, determining activity of E2 EP3 III in presence of regulator and/or compound at varying concentrations.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Sequence 20 BP; 4 A; 4 C; 6 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Example 2; SEQ ID NO 4; 130pp; English.
                                   10-FEB-2004; 2004WO-EP001196
                                                                                              24-FEB-2003; 2003EP-00003253
                                                                                                                                                           (FARB ) BAYER HEALTHCARE AG.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Local Similarity 94.1 tes 16; Conservative
                                                                                                                                                                                                                      Golz S, Brueggemeier U,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (revised)
                                                                                                                                                                                                                                                                                WPI; 2004-653028/63.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      25-MAR-2003
01-JUN-1992
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AAQ21516;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           di sease.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Matches
      8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   The invention relates to treating an inflammatory disorder in an individual comprising administering an oligonucleotide (an antisense coligonucleotide) up to 30 nucleotides in length complementary to a nucleic acid molecule encoding human tumour necrosis factor-alpha (TMF-CC alpha). The oligonucleotide useful in treating an inflammatory disorder comprises at least as 8 nucleobase portion of any of 50 20-21 base pair sequences, given in the specification. The antisense oligonucleotide is administered orally, topically or parenterally. The oligonucleotide is comprises at least one modified intersugar linkage. The intersugar linkage is a phosphorothicate linkage. The oligonucleotide further comprises at least one 2'-O-methoxyethyl modification and at least one 5'-O-methoxyethyl modified cytidine residue is a 5-methyl cytidine, where every cytidine residue is a 5-methyl cytidine, and where every cytidine seature (is a 5-methyl cytidine and the modified intersugar linkage is a methylene(methylimino) cr incrsugar linkage. The methods and compositions of the present invention are useful for the diagnosis, prevention and/or treatment of diseases or conditions associated with aberrant expression or activity of the TNF-CC alpha, such as inflammatory, infectious and autoimmune diseases, contitions and hepatisis, theopression and autoimmune disease, pancreatitis, including diabetes, rheumatoid arthritis, crohn's disease, pancreatitis, colitis and hepatisis. The present sequence is an antisense
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                       Treating inflammatory disorders, such as diabetes, rheumatoid arthritis and multiple sclerosis, using antisense oligonucleotides targeted to nucleic acids encoding human tumor necrosis factor-alpha (TNF-alpha).
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ss; prostaglandin E2 EP3 III; hematological disease; cardiovascular disease; urological disease; metabolic disease; endocrinological disease; gastroenterological disease; cancer; respiratory disease; regulator; primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.4%; Score 15.4; DB 1; Length 20; 94.1%; Pred. No. 2.2e+02; ive 0; Mismatches 1; Indels.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               oligonucleotide targeting the human TwFalpha gene.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 20 BP; 8 A; 3 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                  Butler MM, Shanahan WR;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Example 22; SEQ ID NO 228; 145pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Prostaglandin B2 EP3 III reverse primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  3416 TCAAGGAAGTATGGAAA 3432
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1 TCAAGGAAGTCTGGAAA 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            BP
02-APR-2001; 2001US-00824322
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ADR67866 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             18-NOV-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        16; Conservative
                                                                                                                                                                                                                  Baker BF, Bennett CF,
                                                                                                                       (BUTL/) BUTLER M M.
(SHAN/) SHANAHAN W R.
                                                             BAKER B F.
BENNETT C F.
                                                                                                                                                                                                                                                                             WPI; 2004-580193/56.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADR67866;
                                                          (BENN/)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Best Loca
Matches
```

```
ö
                      Gaps
                      ö
                                                                                                                                                                                                           MDR; chemotherapy; PCR; cytosine triphosphate sythetase; ss
0.4%; Score 15.4; DB 1; Length 20; 94.1%; Pred. No. 2.2e+02; tive 0; Mismatches 1; Indels
                                                                                                                                                                                       Mutant primer for amplifying exon 4 of CTP synthetase gene.
                                                                                                                                                                                                                                                   Location/Qualifiers
                                                                                                                                                                                                                                                                        /*tag= a
/note= "mutation"
                                           2834 TCAAGGAGCTTCCAGTG 2850
                                                                                            RESULT 241
AAQ21516/c
ID AAQ21516 standard; DNA; 20 BP.
                                                          1 TCATGGAGCTTCCAGTG
                                                                                                                                                                   (first entry)
                                                                                                                                                                                                                                                             misc difference 20
                                                                                                                                                                                                                                                                                                       WO9201811-A
                                                                                                                                                                                                                                                                                                                           06-FEB-1992
```

WO2004074830-A2

02-SEP-2004

Bapiens

Homo

ò a Synthetic

10001863-3.81.rng

```
DNA or RNA was purified from white blood cells of patients with acute leukaemia or solid tumour cells; the RNA was reverse transcribed into cDNA. The downstream primer (AAQ21515) was used together with this primer which has a mutant-specific nucleotide at its 3' end to amplify a region of exon 4 of the CTP synthetase gene. A specific signal will only be seen with samples contry. The mutant gene. See AAQ21488 and AAQ21512 for upstream primers which can be used with other primers specific for mutations in exon 4. (Updated on 25-MAR-2003 to correct PA field.)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             DNA encoding cpds. which catalyse de-acetylation - and sulphation of glycosamino-glycan(s), used to develop agonists or antagonists for use in
                                                                                                                                                      Assay method for CTP synthetase mutation(s) - utilising polymerase chain reaction to reveal the presence of the multiple drug resistance phenotype indicative of a mutation.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               N-heparan sulphate-N-deacetylase-N-sulphotransferase; heparan; heparin; blood clotting; glycosaminoglycan; proteoglycans; neoplastic disease; viral infection; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Primer for preparing probe for N-heparan sulphate sulphotransferase.
                                                                                                                                                                                                                                                                                                                                                                                             Score 15.2; DB 1; Length 20;
Pred. No. 2.3e+02;
0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Swiedler S,
                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 6 A; 3 C; 6 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Hashimoto Y,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                MASSACHUSETTS MEDICAL CENT
                                                                     (IMCR ) IMPERIAL CANCER RES TECHNOLOGY.
                                                                                                                                                                                                                                                                                                                                                                                                                                                        1185 AAATTTGGACAGTTTCCCAC 1204
                                                                                                                                                                                                               Example 2; Page 33; 46pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AAACTTGGACAGTTTTCCCC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         BP.
90GB-00016287.
                           90GB-00016287.
                                                                                                                                                                                                                                                                                                                                                                                              0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            93WO-US005412
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        92US-00057167
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Orellana A,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AAQ53908 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                          17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (revised)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 1994-007517/01.
                                                                                                                             WPI; 1992-064967/08
                                                                                                                                                                                                                                                                                                                                                                                                              Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (UYMA-) UNIV MASS
(GLYC-) GLYCOMED
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Hirschberg CB,
Ishihara M;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          07-JUN-1993;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WO9325659-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        16-JUN-1992;
 25-JUL-1990;
                           25-JUL-1990;
                                          11-APR-1991;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    30-APR-1993;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              25-MAR-2003
22-JUN-1994
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 23-DEC-1993
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAQ53908;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20
                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                                                                                  Meuth M;
                                                                                                                                                                                                                                                                                                                                                                                                                             Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT
 ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 셤
```

2 Wei

Gaps

.. 0

```
·.
                                                          N-heparan sulphate sulphotransferase can catalyse the deacetylation and sulphation of a glycosaminoglycan (GAG), namely heparan sulphate. It can be used for treating conditions in which the stimulation or inhibition of the deacetylation and or sulphation of heparan is desirable, such as blood clotting disorders, neoplastic conditions and viral infection. The enzyme can also be used to produce highly modified and therefore highly active sulphated proteoglycans for use in therapy. This primer was used to amplify a subclone of the enzyme coding sequence to produce a probe. (Updated on 25-MAR-2003 to correct PN field.)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Diagnosis of X-linked severe combined immunodeficiency (XSCID) - comprises detecting mutated IL-2R gamma gene, also vectors and transgenic animals containing the mutated gene.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AAQ71911 to AAQ71975 are primers for the human IL-2R gammma gene, these were used to amplify DNA from mutated and normal IL-2R gamma genes. The mutated gene DNA was obtained either from female carriers or male sufferers of X-linked severe combined immunodeficiency (XSCID). The amplified DNA from normal and affected individuals was then compared using a variety of methods including northern blotting and dot and slot hybridistation. From this a claimed method for the diagnosis of XSCID carriers and sufferers was developed. (Updated on 25-MAR-2003 to correct PN field.)
                                                                                                                                                                                                                                                                                                       Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    IL2-R gamma gene; X-linked severe combined immunodeficiency; XSCID;
interleukin; ss.
                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                  0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; ive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Sequence 20 BP; 4 A; 4 C; 4 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                     Seguence 20 BP; 0 A; 4 C; 7 G; 9 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Human IL-2R gamma gene exon 4 seq 2 primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (USSH ) US DEPT HEALTH & HUMAN SERVICES.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Mcbride WO;
                                Disclosure; Page 23; 72pp; English.
                                                                                                                                                                                                                                                                                                                                         1840 CCAAAAACAGGAACTACAG 1859
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Claim 12; Page 88; 98pp; English.
                                                                                                                                                                                                                                                                                                                                                                   20 CCAGAAAACAGGCACCACAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                            BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       94WO-US002891.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         93US-00031143
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       93US-00121435
                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAQ71950 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (first entry)
                                                                                                                                                                                                                                                                                       l Similarity 85.0
17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Noguchi M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (revised)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           WPI; 1994-303046/37.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WO9420641-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       10-MAR-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         12-MAR-1993;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         14-SEP-1993;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Leonard WJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              25-MAR-2003
03-MAY-1995
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     15-SEP-1994.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AAQ71950;
                                                                                                                                                                                                                                                                         Query Match
                                                                                                                                                                                                                                                                                          Local
therapy
                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 243
                                                                                                                                                                                                                                                                                                          Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                            AAQ71950
g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
```

10001863-3.81.rng

Capped RNA molecule; mRNA maturation; translation initiation; influenza; endonuclease aptamer; RNase; therapy; inhibitor; ss.

Synthetic

5' fragment #2 of alfalfa mosaic virus.

(first entry)

27-AUG-1997

AAT47265;

AAT47265 standard; RNA; 20 BP.

```
AAT47265/
                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The method aims to provide a collection of highly reproducible microsatellite marker sequences (MMS) at approx. 10-50 cM intervals throughout the human genome which can be detectably labelled. The MMS are polymorphic, simple sequence repeats and can be used in automated genotyping. esp. fluorescence-based. The primers correspond to the unique DNA sequence surrounding each marker, and PCR is used to detect each polymorphism. When the MMS show considerable polymorphism (ie. a difference in the number of repeats) between individuals, the markers can be particularly informative. The MMS can be ideal for linkage studies. Kits comprise at least 4 groups, of at least 3 sets, each comprising are shown in AAQ95687.734. The published size range of the DNS. Is allele as the control of the DNA. Group 7 primer pairs are shown in AAQ95687.734. The published size range of the DiSSII8 allele is a 118-120 bp, and the degree of heterozygosity in the population is
                                                                                                                                                                                                                                                                                                                                                                                                                                                   Kit for automated genotyping contg. pairs of PCR primers - designed to amplify polymorphic nucleotide repeat sequences, arranged in sets each with a characteristic fluorescence label, useful e.g. in detection of disease related genetic rearrangement.
                       Gaps
                       ;
0
                                                                                                                                                                                                                 primer; polymerase chain reaction; PCR; linkage study; locus; microsatellite marker sequence; automated genotyping; allele; polymorphism; detection; Homo sapiens; ss.
                                                                                                                                                                                             Primer A (Group 7, set C) for marker D15S118, chromosome 15.
  Length 20;
                      3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 20 BP; 9 A; 7 C; 1 G; 3 T; 0 U; 0 Other;
Disclosure, Fig 7G-2; 104pp; English.
                                         2093 AGTATCTGTTGTAGCAGTTC 2112
                                                                1 AGAATCTGTTGTTCCAGTTC 20
                                                                                                                                                                                                                                                                                                                                     94WO-US013945.
                                                                                                                                                                                                                                                                                                                                                           93US-00160837.
                                                                                                                          AAQ95706 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                SNINGO NING (OCYU)
                                                                                                                                                                      (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                           WPI; 1995-215278/28.
         Local Similarity
                                                                                                                                                                                                                                                                                                                                     05-DEC-1994;
                                                                                                                                                                                                                                                                                                                                                           03-DEC-1993;
                                                                                                                                                                      15-FEB-1996
                                                                                                                                                                                                                                                                                        WO9515400-A1
                                                                                                                                                                                                                                                                                                              08-JUN-1995
                                                                                                                                                                                                                                                                                                                                                                                                        Levitt RC;
                                                                                                                                                                                                                                                                 Synthetic.
                                                                                                                                                AAQ95706;
 Query Match
                                                                                                    RESULT 244
                     Matches
                                                              g
```

```
AAT47264-T47280 represent capped RNA molecules produced by the method of the invention. The method of the invention is for producing capped RNA or RNA analogues. The method comprises reacting a RNA or analogue oligonucleotide with a phosphate addition agent to form a RNA or analogue oligonucleotide with a phosphate addition agent to form a RNA or analogue mono-, di- or triphosphate, which is then capped. The presence of the cap is important for mRNA maturation, initiation of translation, and protects the mRNA against various RNases present in the cell. The capped RNA or analogue is an influenza endonuclease aptamer, useful for treating or preventing an influenza infection in an animal. The synthetic capped RNA are substrates for virally encoded endonuclease associated with influenza virus. The short non-extendible (due to their length or because of the modification of the 3' end of the oligo) RNA molecules are potent inhibitors of the clawayse of capped RNA by influenza endonuclease. They may be used to investigate viral and cellular mechanisms of transcription/translation, or mRNA maturation
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Production of capped RNA or analogues - useful as substrates for influenza virus associated virally encoded endonuclease.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 2.3e+02; Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 20 BP; 3 A; 1 C; 2 G; 0 T; 14 U; 0 Other;
                                                                                                                                                                                                                                                                                                                /mod_base= 2'-0-methyluridine
                                                                    *tag= a
|mod_base= 7-methylguanosine
                                                                                                                                                               /*tag= b
/mod_base= triphosphorylated
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Olsen DB
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   GAAGAGGAAAATAAAAACC 2702
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Claim 18; Page 12; 39pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    20 GAAAATTAAAAATAAAACC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Kuo LC,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              96WO-US008394.
                                                                                                                                                                                                                                                                               *tag= c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (MERI ) MERCK & CO INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Benseler F, Cole JL,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 1997-051868/05.
                                modified base
                                                                                                                                     modified_base
                                                                                                                                                                                                                                           modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              03-JUN-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            07-JUN-1995;
                                                                                                                                                                                                                                                                                                                                                                                     WO9640159-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                         19-DEC-1996
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2683
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  · · Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ò
```

RESULT 246

ö

Gaps

ö

0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; tive 0; Mismatches 3; Indels

3389 TCAAATATCCATATTAACCA 3408

ઠ

17; Conservative

Best Local Similarity Matches 17; Conserv

Query Match

1 TCAAAGACCCATATCAACCA 20

AAT89739

```
The primers AAV59158-V59159 were used to detect a point mutation in a p53 agene as an example of the method of the invention to Detect mutant alleles among an excess of wild-type alleles. This is carried out by separating the wild type alleles using a carrier on which are bound one or more oligonucleotides complementary to the wild type. The method is invest to detect mutants having one or more point mutations, deletions, inversions, insertions and/or substitutions of small arge genetic regions, particularly to detect genetic abnormalities for cancer diagnosis. The method can detect cancer cells in faeces, sputum, bronchial lavage, urine or tissue biopsies. Rare alleles can be detected, making the method suitable for detecting heterozygous or homozygous mutations or polymorphisms of any origin
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Vaccine, eye disease; conventional trachoma; nonendemic trachoma; paratrachoma; inclusion conjunctivitis; genital disease; perihepatitis; nongonococcal uretritis; epidymitis; cervicitis; salpingitis; PCR primer; bartholinitis; pneumopathy; venereal lymphogranulomatosis; ss.
                                                                                                Detecting mutated alleles in presence of excess wild-type allele - by separation of wild-type on support carrying specific oligonucleotides, used to detect mutation(s), particularly for cancer diagnosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        PCR primer used to amplify an ORF of Chlamydia trachomatis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
.ive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 2 A; 8 C; 1 G; 9 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Genome sequence of Chlamydia trachomatis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Disclosure; Page 1543; 1755pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               3173 TTTTAAGTCTGTCTCTTAC 3192
                                                                                                                                                                         Example 2; Page 10; 17pp; German.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1 rrrchacrcrercrercrec 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          97FR-00015041.
97FR-00016034.
98US-0107077P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           662/c
AAZ02662 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           98WO-IB001939.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Chlamydia trachomatis
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 1999-371125/31.
                                                               WPI; 1998-495865/42
 ပ
(WAGE/) WAGENER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (GEST ) GENSET
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WO9928475-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           27-NOV-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             28-NOV-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            17-DEC-1997;
04-NOV-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           07-OCT-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          10-JUN-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Griffais R;
                                  Wagener C;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAZ02662;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 248
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AAZ02662,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AAT89689-T89744 are individually claimed oligonucleotides used as PCR (polymerase chain reaction) primers for the discrimination of the genotype of hepatitis C virus (HCV). Classification of the genotype of HCV can be achieved precisely and simply according to the International Standardisation of Classification. The primers can be used to distinguish between HCV genotypes la, lb, 2a, 2b, 3a, 3b, 4, 5a, 6a and 6b
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Primers used for determining hepatitis C virus genotype - provide a rapid and accurate method of hepatitis C virus genotyping.
                                                                                                                                                         Hepatitis C virus; HCV; genotype determination; 1a; 1b; 2a; 2b; 3a; 3b;
4; 5a; 6a; 6b; diagnosis; amplification; PCR; primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             .;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              88; PCR; primer; amplification; mutant allele; carrier; cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Score 15.2; DB 1; Length 20;
Pred. No. 2.3e+02;
0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 20 BP; 2 A; 7 C; 7 G; 4 T; 0 U; 0 Other;
                                                                                                                        primer used for hepatitis C virus genotyping.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                3762 CAGATGGCTGGGATCCCTCC 3781
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Claim 51; Page 18; 33pp; Japanese.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               cacerederiedearcecree 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        98WO-DE000676.
                                                                                                                                                                                                                                                                                                                                  96JP-00038875.
                                                                                                                                                                                                                                                                                                                                                                   95JP-00035997.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       97DE-01008758
                AAT89739 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AAV59158 standard; DNA; 20
                                                                                   (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          14-DEC-1998 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Local Similarity 85.0
nes 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 1997-497313/46.
                                                                                                                                                                                                            Synthetic.
Hepatitis C virus.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             p53 sense primer.
                                                                                                                                                                                                                                                                                                                                                                                                                        (SRLS-) SRL KK
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       04-MAR-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     04-MAR-1998;
                                                                                                                                                                                                                                                               JP09234072-A
                                                                                                                                                                                                                                                                                                                                                                     01-FEB-1995;
30-DEC-1995;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WO9839472-A2
                                                                                                                                                                                                                                                                                                                                  01-FEB-1996;
                                                                                     05-FEB-1998
                                                                                                                                                                                                                                                                                                  09-SEP-1997.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      11-SEP-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Synthetic.
                                                   AAT89739;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AAV59158;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 247
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    PCR
```

ð 셤

ö

Gaps

; 0

```
PCR primers AAZ01426-Z06209 were used to amplify open reading frames (ORFs) of the genome of Chlamydia trachomatis (see AAZ01425). These ORFs encode polypeptides (see AAY36754-Y37949) which can be used as vaccines against Chlamydia trachomatis. Antisense and ribozyme sequences can also be used to control growth of the microorganism. Chlamydia trachomatis is responsible for a large number of diseases, e.g. eye diseases such as conventional trachoma, nonendemic trachoma, paratrachoma, and inclusion conjunctivitis; genital diseases such as nongonococcal uretritis, preumopathy in breast feeding infants; and venereal lymphogranulomatosis. The polypeptides of the invention may be of use in treating these
                                                                                                                                                                                                                                                                                                                                                                                                       Sequence 20 BP; 6 A; 6 C; 3 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                              diseases
X88888888888888888888888888888888888
```

ö Length 20; 3; Indels 0.4%; Score 15.2; DB 1; 85.0%; Pred. No. 2.3e+02; 0; Mismatches 3186 TCCTTACAGAGGTTAAAGTC 3205 20 rrcrrgcagaggrraaaggc 1 Local Similarity 85.0 nes 17; Conservative Query Match Matches

셤

ô

Gaps

AAX92717 standard; DNA; 20 13-SEP-1999 (first entry) AAX92717; 

Respiratory disease; pneumonia; bronchitis; heart disease; sarcoidosis; sinusitis; purulent otitis media; erythema nodosum; pharyngitis; vaccine; neutralising epitope; PCR primer; 88. PCR primer used to amplify an ORF of Chlamydia pneumoniae.

Chlamydophila pneumoniae. Synthetic.

WO9927105-A2

98WO-IB001890 20-NOV-1998; 03-JUN-1999

Antisense compounds useful for treating or preventing infection, inflammation or tumor formation by inhibiting expression of human MEKKS.

3

Gaarde

Cowsert LM,

Monia BP,

WPI; 2000-464034/40.

(ISIS-) ISIS PHARM INC

Sequences AAA61956-A61995 represent phosphorothioate antisense

Claim 3; Col 39; 35pp; English.

97FR-00014673. 21-NOV-1997; 04-NOV-1998;

(GEST ) GENSET

Griffais R;

WPI; 1999-357842/30

Genome sequence of Chlamydia pneumoniae.

Page 1533; Disclosure; 1912pp; English.

Sequences AAAA195-ABOR Expessent phosphorocinoate antisense coligonucleotides targetted to the human MEKES gene, which inhibit its expression. The antisense oligonucleotides were designed to target different regions of the human MEKES RNA, and were analysed for their cffect on MEKES manA levels by quantitative real-time PCR. MEKES (also known as mitogen-activated protein kinase kinase 5, MEK kinase 5, apoptosis signal-regulating kinase 1, and ASK1) is a dual-specific serine/threonine kinase which mediates cellular ceminal kinase/stress-activated protein kinase) and p38 modules of MAP KN cerminal kinase activated protein kinase) and p38 modules of MAP kinase cascades. MEKES is thought to play a critical role in the cegnistion of apoptosis (programmed cell death) by interacting with other regulation of apoptosis (programmed cell death) by interacting with other as MKK3 and SEKI. MEKK5 is proteined of mornistiant of MEKK5 is induced by tumour necrosis factor as MKX3 and SEKI. MEKK5 is participates in another apoptosis-related signalling cascade involving the modulation of transcription factors. Activation and dimerisation of MEKK5 is induced by tumour necrosis factor—alpha (TNF-alpha), these processes being mediated by recoiles of the invention and dimerisated with MEKK5 and inhibit MEKK5 conditions associated with MEKK5 expression, such as inflammation and wound healing disorders

AAX91991-X97517 represent PCR primers used to amplify open reading frames and other nucleic acid sequences from the genome of Chlamydia pneumoniae (see AAX91990). C. pneumoniae causes respiratory disease such as pneumonia and bronchitis and is thought to be a contributing factor in heart disease, sarcoidosis, sinusitis, purulent citis media, erythema nodosum or pharyngitis. The polypeptides encoded by the open reading frames of the C. pneumoniae genome (see AAX34844- AAX3879) can be used in immunogenic compositions as vaccines. Vectors containing C. pneumoniae mucleotides sequences can also be used as immunogenic compositions, especially where the vector directs the expression of a neutralising epitope of C. pneumoniae

.

```
ö
                                                                                                                                                                                                                    MEK kinase 5; MAP/ERK kinase kinase 5; ASK1; pro-apoptotic; apoptosis signal-regulating kinase 1; programmed cell death; serine/threonine kinase; MAP kinase cascade; JMK/SAPK module; JUN N-terminal kinase/stress-activated protein kinase; p38 module; MKK3; SEK1; transcription factor modulation; expression inhibition; antisense; inflammation; wound healing disorder; phosphorothioate; ss.
                                                                                                                                                                                         Human MEKKS phosphorothioate antisense oligonucleotide, SEQ ID NO:31.
                                       Gaps
                                       ö
                                                                                                                                                                                                            Human MEXXS; mitogen-activated protein kinase kinase kinase
                  0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.38+02;
iive 0; Mismatches 3; Indels
Sequence 20 BP; 5 A; 2 C; 7 G; 6 T; 0 U; 0 Other;
                                                          1617 GAAAACTTCCTTCCAGATAT 1636
                                                                      20 GAAACTTCCCTCCAGTCAT 1
                                                                                                                               AAA61979 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                      99US-00359757.
                                                                                                                                                                                                                                                                                                                                                                        99US-00359757.
                                                                                                                                                                     (first entry)
                                      17; Conservative
                           Local Similarity
                                                                                                                                                                                                                                                                                                                                                      23-JUL-1999;
                                                                                                                                                                                                                                                                                            Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                        23-JUL-1999;
                                                                                                                                                                     20-NOV-2000
                                                                                                                                                                                                                                                                                                                                  27-JUN-2000.
                                                                                                                                                  AAA61979;
                  Query Match
                                      Matches
                                                                                                           RESULT 250
                                                                                                                      AAA61979,
                                                                                                                               셤
```

S

RESULT 251

ð 셤

```
Novel compound for diagnosing, preventing and treating immune disorders, comprising an oligonucleotide that specifically hybridizes with a nucleic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            The present invention provides sequences of antisense oligonucleotides targeted at the murine and human B7-1 and B7-2 coding and mRNA sequences. The antisense sequences have phosphorothicate backbones and some nucleotides are 2'-methoxyethoxy residues. The sequences can be used in the treatment of inflammatory and autoimmune disorders, including asthma, juvenile diabetes mellitus, myasthenia gravis, Graves' disease, rheumatoid arthritis, allograft rejection, inflammatory bowel disease, multiple sclerosis, psoriasis, systemic lupus erythematosus, contact dermatitis, rhinitis, allergies and cancer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human; ATM; ataxia telangiectasia; mutation detection;
single-stranded conformation polymorphism; SSCP; electrophoresis;
                                                                       Human, mouse; B7-1; B7-2; antisense; PCR primer; inflammation; autoimmune disorder; phosphorothioate backbone; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          / Match 0.4%; Score 15.2; DB 1; Length 20; Local Similarity 85.0%; Pred. No. 2.3e+02; nes 17; Conservative 0; Mismatches 3; Indels
                                   Human B7-1 antisense oligonucleotide SEQ ID NO: 254.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 7 A; 4 C; 8 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human ATM gene exon 40 reverse primer.
                                                                                                                                                                                                                                                                                                                                                                                    Karras JG;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Example 19; Page 98; 162pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                3767 GGCTGGGATCCCTCCCTGT 3786
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          acid sequence encoding B7 protein.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    20 decredenticereretri 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AAF60191 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       21-JUL-2000; 2000WO-US020011.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 99US-00360416.
                                                                                                                                                                                                                                                              25-MAY-2000; 2000WO-US014471.
                                                                                                                                                                                                                                                                                                      99US-00326186.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (first entry)
(first entry)
                                                                                                                                                                                                                                                                                                                                                                                    Vickers TA,
                                                                                                                                                                                                                                                                                                                                              (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2001-049991/06.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WO200107660-A1
                                                                                                                                                                                WO200074687-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 23-JUL-1999;
                                                                                                                                                                                                                                                                                                      04-JUN-1999;
23-MAR-2001
                                                                                                                                           Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           27-APR-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   01-FEB-2001.
                                                                                                                                                                                                                                                                                                                                                                                    Bennett CF,
                                                                                                                                                                                                                        14-DEC-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAF60191;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 253
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAF60191
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0;
                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   The present invention provides sequences of antisense oligonucleotides targeted at the murine and human B7-1 and B7-2 coding and mRNA sequences. The antisense sequences have phosphorothioate backbones and some nucleotides are 2'-methoxyethoxy residues. The sequences can be used in the treatment of inflammatory and autoimmune disorders, including asthma, juvenile diabetee mellitus, myathenia gravis, Graves' disease, rheumatoid arthritis, allograft rejection, inflammatory bowel disease, multiple sclerosis, psoriasis, systemic lupus erythematosus, contact
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Novel compound for diagnosing, preventing and treating immune disorders, comprising an oligonucleotide that specifically hybridizes with a nucleic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ;
0
                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                            Human, mouse, B7-1; B7-2; antisense; PCR primer; inflammation; autoimmune disorder; phosphorothioate backbone; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
0.4%; Score 15.2; DB 1; Length 20;
Best Local Similarity 85.0%; Pred. No. 2.3e+02;
Matches 17; Conservative 0; Mismatches 3; Indels
                                 0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
tive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                         Human B7-1 antisense oligonucleotide SEQ ID NO: 128.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 20 BP; 3 A; 4 C; 4 G; 9 T; 0 U; 0 Other;
  Sequence 20 BP; 5 A; 6 C; 2 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       dermatitis, rhinitis, allergies and cancer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Karras JG;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Example 12; Page 75; 162pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1317 TTGAGTTTCAAAGGTTGCTG 1336
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        comprising an oligonucleotide that acid sequence encoding B7 protein.
                                                                                                                        2236 AGCTAGTAAAGAATTTAGAA 2255
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         rrradrircacadcriderg 20
                                                                                                                                                           20 AGCTGGTAGAGACTTTAGAA 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAF33171 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                     ВЪ.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           25-MAY-2000; 2000WO-US014471.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     99US-00326186
                                                                                                                                                                                                                                                                     AAF32931 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                   23-MAR-2001 (first entry)
                                 V.**
Dest Local Similarity 85.09
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Bennett CF, Vickers TA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 2001-049991/06
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WO200074687-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     04-JUN-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       14-DEC-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AAF33171;
                                                                                                                                                                                                                                                                                                           AAF32931;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 252
AAF33171/C
ID AAF331:
XX
AC AAF331:
XX
```

AAF32931
AAF

유 ò

ö

Gaps

ö

```
The present sequence is one of a number of primers used in a method for detecting a mutation or a polymorphism in the human ATM gene, which is associated with the disease ataxia telangiectesia, or a polymorphism in the human ATM gene, which is casociated with the disease ataxia telangiectesia.

Single-stranded conformation polymorphism (SSCP) electrophoresis that allows electrophoresis of two or three amplified segments in a single lane. The method is useful for screening large, complex polyworphisms. The method is useful for screening large, complex polyworphisms. The method is useful for screening of human DNA samples for mutations, for distinguishing mutations from polymorphisms, and for improving the efficiency of automated screening method. The mega-SSCP method provides a screening method of genes for multiple polymorphisms and mutations at once. The method is particularly suitable for large, polyexonic, cukaryotic genes, having mutations and polymorphisms at many points and not merely at one or a few hot spots. Note: the SEC ID assigned to this sequence in the disclosure and claims of the the specification is one
                                                                                                                                             Detecting a mutation or polymorphism in human ataxia telangiectasia gene or polyexonic eukaryotic gene, involves using mega-single stranded comformation polymorphism analysis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human nucleolin phosphorothioate antisense oligonucleotide, SEQ ID NO:51.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human nucleolin; P92; C23; phosphoprotein; ribosome biogenesis; ribosome transport; cytokinesis; nucleogenesis; cell proliferation; cell growth; transcriptional repression; replication; esignal transduction; chromatin decondensation; Ag-NOR aucleolin antibody; systemic connective tissue disease; SLE; systemic lupus erythematosus; celeroderma-like chronic graft versus host disease; scleroderma-like chronic graft versus host disease; expression inhibition; tumour formation; cancer; inflammation; immune disorder; phosphorothioate; antisense oligonucleotide; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     number lower than the number given in the sequence listing
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 20 BP; 6 A; 7 C; 1 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1989 TGTGCAACACCTTCAGATAA 2008
                                                                                                                                                                                                                                    Claim 7; Page 54; 118pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1 TTTGCAACACCTTCACCTAA 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AAC92601 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   27-MAR-2001 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Best Local Similarity 85.0
Matches 17; Conservative
                    (REGC ) UNIV CALIFORNIA
                                                                                                    WPI; 2001-168574/17.
                                                            Gatti RA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AAC92601;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      RESULT 254
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAC92601/
ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ద
```

```
Sequences AAC92560-C92639 represent antisense oligonucleotides targetted to the human nucleolin gene, which inhibit its expression. The antisense coligonucleotides were designed to target different regions of the human nucleolin mRNA and were analysed for their effect on nucleolin mRNA levels by quantitative real-time PCR. Nucleolin (also known as P92 or 10.3) is the most abundant nucleolar phosphoprotein in actively growing cells. Nucleolin primarily participates in ribosome biogenesis and transport of ribosomeal components, being able to transfently bind to pretribosomes in the nucleolus via a ribonucleoprotein consensus sequence. However, it has also been shown to be involved in cytokinesis, nucleogenesis, cell proliferation and growth, transcriptional repression, replication, signal transduction, and chromatin decondensation. Nucleolin is a member of the Ag-NOR (active ribosomal gene located in the nucleolar or ganiser region) family of proteins which are markers of active or tumour growth rate. The presence of antibodies against nucleolin are associated with systemic connective tissue diseases such as systemic lupus erythematosus (SLE) and scleroderma-like chronic graft versus host lupus erythematosus (SLE) and scleroderma-like chronic graft versus host greavention and treatment of conditions associated with nucleolin expression, such as tumour formation, immune disorders and inflammation
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  domain;
                                                                                                               Novel antisense compound targeted to human nucleolin which specifically hybridizes with and inhibits the expression of human nucleolin, useful for modulating the expression of nucleolin in cells.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Antisense gene therapy, RAIDD; death domain, caspase recruitment de CARD, hyperproliferative disorder, cancer; growth disorder, mouse, metabolic disorder, infection; inflammation; tumour formation; RIP associated ICH-1/CBD-3-homologous protein with death domain; receptor interacting protein; antisense oligonucleotide; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.4%; Score 15.2; DB 1; Length 20; 35.0%; Pred. No. 2.3e+02; ve 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 7 A; 2 C; 5 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Mouse RAIDD antisense oligonucleotide #52.
                                                                                                                                                                                                     Claim 14; Col 43-44; 41pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               978 TTTGACAAATCTGCTCTAGA 997
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20 rrraacaaarcrecrea 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ABK99798 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 29-OCT-2001; 2001WO-US050914.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          01-NOV-2000; 2000US-00705267.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   85.0%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Local Similarity 85.0 ses 17; Conservative
                                      Cowsert LM;
(ISIS-) ISIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (ISIS-) ISIS PHARM INC
                                                                               WPI; 2001-079848/09.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WO200248314-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Mus musculus.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 21-OCT-2002
                                        Bennett CF,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20-JUN-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ABK99798;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 255
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ABK99798/
  ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      셤
```

ö

Gaps

ö

Score 15.2; DB 1; Length 20; Pred. No. 2.38+02; 0; Mismatches 3; Indels

0.4%;

99US-00433699. 99US-00433699.

03-NOV-1999; 03-NOV-1999;

Homo sapiens

US6165786-A.

```
Dna marker associated with obesity, used in a predictive method of
likelihood of obesity.
                                                                                                                                                                                                                                                                   Example 2; Page 5; 8pp; Korean.
                                                                                                                                                                                                                                                 WPI; 2002-737873/80.
       WPI; 2002-583496/62
                                                                                                                                                                                                                                         Kim GT,
                                                                                                                                                                                                     KR2002041152-A.
                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                       04-MAR-2003
                                                                                                                                                                                                            01-JUN-2002
                                                                                                                                489
                                                                                                                                      20
                                                                                                                                                                                                                                         Kang BY,
                                                                                                                                                                ABX14629;
                          disorder
                                                                                                                 Query Match
Zhang H,
                                                                                                                                                  RESULT 256
                                                                                                                       Matches
                                                                                                                                                     ABX14629
ò
                                                                                                                                     ద
```

Cauet G, 2933 GAAAGICATIICAACICITA 2952 Disclosure; Page 78; 79pp; French. 1 gacagrerrireacererra 20 882/c ABT11882 standard; DNA; 20 BP. 29-JAN-2002; 2002WO-FR000348. 31-JAN-2001; 2001FR-00001294. Achstetter T, (first entry) (AVET ) AVENTIS PHARMA SA. 17; Conservative Query Match Best Local Similarity WPI; 2002-723143/78. WO200261109-A2 Unidentified Spagnoli R, 19-DEC-2002 08-AUG-2002. Winter J; ABT11882; Matches ABT11882 RESULT 888888888888888888888 ð 셤 The invention describes a compound (I) 8-50 nucleobases in length
targeted to a nucleic acid molecule (II) encoding RAIDD which is an
adaptor molecule containing both death domain (DD) and caspase
c recruitment domains (CARD), where (I) specifically hybridises with and
c inhibits expression of RAIDD, or specifically hybridises with at least an
c inhibiting the expression of RAIDD (Receptor interacting protein (RIP)
c associated (CH-1/CED-3-homologous protein with death domain) in cells or
tissues, and for treating an animal having a disease or condition
c associated with RAIDD, where the disease or condition is a
hyperproliferative disorder such as cancer, or a growth or metabolic
disorder. (I) is also useful for distinguishing functions of various
as research reagents and kits, for distinguishing functions of various
c an inhibitation or tumour formation. This sequence represents a mouse RAIDD
c antisense oligonucleotide used to control expression of the RAIDD protein ° Novel antisense compound that hybridizes and inhibits nucleic acid encoding RAIDD which is an adaptor molecule containing both death domain and caspase recruitment domains, for treating hyperproliferative Gaps 6 0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; ive 0; Mismatches 3; Indels Human; 88; PCR; primer; lipoprotein lipase; LPL; obesity. Sequence 20 BP; 4 A; 2 C; 5 G; 9 T; 0 U; 0 Other; Human lipoprotein lipase PCR primer #2. TTGACAGGAAACCCCATCCA 508 Claim 3; Page 95; 144pp; English Trgacagaaaccacatrca 1 (SEOU-) SEOULIN BIO SCI CO LTD. 27-NOV-2000; 2000KR-00070924. 27-NOV-2000; 2000KR-00070924. ABX14629 standard; DNA; 20 (first entry) Local Similarity 85.0 nes 17; Conservative Lee Freier SM,

The invention relates to a DNA marker associated with obesity and a predictive method of the likelihood of obesity using the DNA marker are provided, used in inhibiting obesity effectively in preventive medicine. The DNA marker associated with obesity has the nucleotide sequence of human LPL (lipoprotein lipase) wherein a 6th base from the origin (+1) of exon 4 in intron 3 of the human LPL is changed from C to T, wherein the base-substituted nucleotide sequence of human LPL comprises a C type allele gene with a MboII restriction enzyme-recognition site, i.e. GAAGA, and a T type allele gene without the MboII restriction enzyme-recognition site. The predictive method of likelihood of obesity comprises the steps of: collecting DNA from a human blood sample; amplifying a region of the intron 3 of the LPL gene in a DNA sample containing a region from -6 to intron 3 of the origin of the exon 4 by PCR; digesting the amplified DNA with MboII; and determining the gene type of the blood sample based on the band size of a digested DNA fragment. The present sequence is a PCR primer used in the above method ö Genetically modified yeast strain, autonomous; metabolism of cholesterol; 17alpha-hydroxypregnenolone; steroid; cortisol; cortexolone; 17alpha-hydroxyprogesterone; ds. New genetically modified yeast, useful for producing therapeutic steroids from simple carbon source, provide high yields at low cost. The invention relates to a genetically modified yeast strain that produces, autonomously from a simple carbon source, a steroid, or its derivative, formed by metabolism of cholesterol. The steroid is 17alpha-hydroxypregnenolone; cortisol; cortexolone or 17alpha-thydroxyprogesterone. The genetically modified yeast strain is used to produce therapeutically useful steroids, and can itself be used as pharmaceutical. This polynucleotide sequence represents an oligomucleotide relating to the steroid producing genetically modified yeast strain of the invention Gaps ä Autonomously steroid producing yeast strain related oligo SEQ ID No Pompon ; 0 0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; tive 0; Mismatches 3; Indels Dumas B, Degryse E, Seguence 20 BP; 4 A; 6 C; 2 G; 8 T; 0 U; 0 Other; 

ö

Gaps

ö

Length 20; 3; Indels

0.4%; Score 15.2; DB 1; 85.0%; Pred. No. 2.3e+02; live 0; Mismatches 3;

g

ð

ö

Gaps . 0

Length 20; Indels

Conservative

Query Match Best Local Similarity Matches 17; Conserv

Sequence 20 BP; 4 A; 6 C; 6 G; 4 T; 0 U; 0 Other;

```
Score 15.2; DB 1;
Pred. No. 2.3e+02;
0; Mismatches 3;
Sequence 20 BP; 6 A; 3 C; 7 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Capture oligonucleptide Zip ID#2210 oligo #9
                                                                                                           0; Mismatches
                                                                                                                                                                    1558 GCATCTTCAATGGCTTGTCC 1577
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Example 5; Fig 29; 300pp; English.
                                                                                                                                                                                                                            GCATCTTCAATGGCCTTACC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gerry NP,
                                                                                                                                                                                                                                                                                                                                                                   ABI95123 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (CORR ) CORNELL RES FOUND INC
                                                     0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                14-APR-2000; 2000US-0197271P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       04-APR-2001; 2001WO-US010958.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    16-FEB-2002 (first entry)
                                                                                                           17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Barany F, Zirvi M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2002-034366/04
                                                                                   Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WO200179548-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  25-OCT-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                               ABI95123;
                                                                                                                                                                                                                         20
                                                     Query Match
                                                                                                                                                                                                                                                                                                       ABI95123

ABI95123

XX

ABI9513

XX

ABI9513

XX

XX

BABI9513

BABI951

XX

BABI951

ABI951

ABIP51

ABIP
                                                                                                           Matches
                                                                                                                                                                                                                       g
                                                                                                                                                                    ઠે
```

```
The present invention describes a method (M1) for designing capture oligonucleotide probes (I) for use on a support to which complementary coligonucleotide probes (II) will hybridise with little mismatch, where (I) have melting temperatures within a narrow range. The method is useful for detecting infectious diseases caused by bacterial infectious agents e.g. Salmonella, Listeria monocytogenes and Haemophilus influenza, fungal infectious agents e.g. Cryptococcus neoformans, Candida albicans and Appergillus funigatutus, viruses e.g. T-cell lymphocytotrophis cirus, Ebstrain-Barr virus and polio virus, and parasitic infectious agents e.g. Ebstrain-Barr virus and polio virus, and parasitic infectious agents medinesis. The method is also useful for detecting genetic diseases such as 21 hydroxylase deficiency, Turner Syndrome and obesity defects.

CC betecting cancer involving oncogenes, tumour suppressor genes involved in DNA amplification, replication, recombination or repair, the cancer is specifically associated with a gene selected from BRCA1 gene, p53 gene, human papillomavirus types 16 and 18 and liver cancers. The method is also used for environmental monitoring, forensics and the food and feed industry, detecting comprises scanning (using e.g. a scanning electron microscope and infrared microscope) the support at the food and feed industry, detecting comprises scanning (using a computer) identified ligation to a presence or absence of the target nucleotide sequences. AB182074 to present invantion
                                                                                                                                                Human, K-ras, PCR primer, probe, capture probe, mutation detection, ligase detection reaction; LDR; p53; BRCAL; BRCA2; infectious disease; infection; 21 hydroxylase deficiency; Turner Syndrome, obsity; cancer; oncogene; tumour suppressor; human papillomavirus; forensic; environmental monitoring; food industry; feed industry; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Designing capture oligonucleotide probes for use on a support to which complementary oligonucleotides hybridize with little mismatch.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Kliman R;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Favis R,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         of the present invention
```

```
The invention relates to antisense oligonucleotides (ACF33752-ACF33779) targeted to the human CREB (CAMP response element binding protein) gene, which inhibit its expression. The oligonucleotides were designed to target different regions of human CREB mENA, and were analysed for their effect on CREB expression by quantitative real-time PCR. CREB (also known as CREB1) is a member of the basic leucine zipper (bZIP) family of transcription factors, and activates transcription of target genes in response to a diverse array of stimuli including peptide hormones, growth factors and protein kinases. It is a component of intracellular signalling events which regulate a wide variety of biological functions,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        New antisense oligonucleotides for modulating CREB (CAMP response elemen
binding protein) gene expression, useful for preventing or treating e.g.
cancers, a disease arising from aberrant apoptosis, or neuronal
                                                                                                                                                                                                                                                                                                                                                    Human; CREB; cAMP response element binding protein; CREB1; bZIP; basic leucine zipper; transcription factor; intracellular signalling; spermatogenesis; circadian rhythm; memory; apoptosis; hyperproliferative disorder; cancer; tumour; blood; soft tissue; apoptosis related disease; neuronal disorder; chromosome 2q32.3-34; cytostatic; neuroprotective; expression inhibition; phosphorothioate;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   backbone with all cytosine residues being 5-methylcytosines. Optionally, it also has 2-methyoxyethyl (2'-MOE) wings at the 5' and 3' ends, which are 5 nucleotides in length"
                                                                                                                                                                                                                                                                                Human CREB phosphorothioate antisense oligonucleotide, SEQ ID NO:21.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /*tag= a
/mod_base= OTHER
/note= "This oligonuclectide has a phosphorothioate
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Claim 3; Page 74; 91pp; English.
763/c
ACF33763 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       antisense oligonucleotide, ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             07-OCT-2002; 2002WO-US032181.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              10-OCT-2001; 2001US-00973827.
                                                                                                                                                                                                   (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (ISIS-) ISIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Cowsert LM;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2003-381663/36.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WO2003030617-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          modified base
                                                                                                                                                                                               24-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                17-APR-2003.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        disorders.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Monia BP,
                                                                                                                  ACF33763;
```

```
including apermatogenesis, circadian rhythms and memory. Overexpression of CREB has been found to induce apoptosis in certain cells, although CREB overexpression may also be linked to cancer as it is constitutively activated in human somatotroph adenomas. CREB may also play a role in the development of drug dependency, as it has been found to mediate morphine-induced upregulation of the CAMP pathways that contribute to opiate dependency. The oligonucleotides of the invention are useful for diagnosis, prevention and treatment of CREB-related disorders, such as hyperproliferative disorders (particularly cancer, e.g., those of blood and soft tissue), diseases or conditions arising from aberrant apoptosis, or neuronal disorders. The present sequence represents a human H-ras phosphorothicate antisense oligonucleotide used as a positive control in determining optimal oligonucleotide concentration for a particular cell
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              for treating or preventing cancer or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /mcd_base= OTHER
/mcd_base= OTHER
/note= "This oligonucleotide has a phosphorothioate
backbone and 2-'methyoxyethyl (2'-MOE) wings at the 5'
and 3' ends, which are 5 nucleotides in length. Also all
cytidine residues are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              New antisense oligonucleotides for modulating vitamin D nuclear receptor gene expression, particularly useful for treating or preventing cancer or
                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         77.
                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               disorder, or as diagnostics or research reagents.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Vitamin D nuclear receptor antisense oligonucleotide, SEQ ID
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Cytostatic; gene therapy; antisense oligonucleotide; human; vitamin D nuclear receptor; cancer; developmental disorder;
                                                                                                                                                                                                                                                                              Length 20;
                                                                                                                                                                                                                                                                                                              3; Indels
                                                                                                                                                                                                                                              Sequence 20 BP; 3 A; 2 C; 10 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                            Score 15.2; DB 1;
Pred. No. 2.3e+02;
                                                                                                                                                                                                                                                                                                            0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    location/Qualifiers
                                                                                                                                                                                                                                                                                                                                              CCCAGCACTTCATCCAGAGC 2396
                                                                                                                                                                                                                                                                                                                                                                           cccascacriccracacase 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Roach MP;
                                                                                                                                                                                                                                                                            0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 13-NOV-2002; 2002WO-US036692.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 14-NOV-2001; 2001US-0000213
                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADB99938 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ø
                                                                                                                                                                                                                                                                                                              Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        l. .20
/*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        phosphorothioate; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Dobie K,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WPI; 2003-468578/44
                                                                                                                                                                                                                                                                                            Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WO2003041657-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 developmental
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           04-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   22-MAY-2003
                                                                                                                                                                                                                                                                                                              17;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Baker BF,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Synthetic.
                                                                                                                                                                                                                                                                                                                                                                            20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADB99938;
                                                                                                                                                                                                                                                                                                                                              2377
                                                                                                                                                                                                                                                                            Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                             RESULT 260
                                                                                                                                                                                                                  line
                                                                                                                                                                                                                                                                                                              Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                            ADB99938
   ઠ
                                                                                                                                                                                                                                                                                                                                                                       g
```

The present invention relates to novel antisense oligonucleotides (ADB99875-ADB99952) which are targeted to a human vitamin D nuclear

Example 15; SEQ ID NO 77; 122pp; English.

ö

```
ó
receptor coding sequence (ADB99864), and specifically hybridizes with and inhibits the expression of vitemin D nuclear receptor. The antisense oligonucleotides are useful for treating an animal having a disease or condition associated with vitamin D nuclear receptor, e.g. cancer or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ហ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     The compound
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   \mathtt{not} \overline{\mathbf{e}} = "Phosphorothioate backbone and all cytosines are -methyl cytosines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    The invention relates to a compound 8-50 nucleobases in length targeted to a nucleic acid molecule encoding complement component C3. The compour specifically hybridises with the nucleic acid molecule encoding
                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                 Antinsense oligonucleotide targeting mouse C3 component, ISIS140094.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   New antisense oligonucleotides targeted to a nucleic acid molecule encoding complement component C3, useful for treating a disease or condition associated with complement component C3, e.g. autoimmune
                                                                                                                                                                                                                                                                                                                                                                                         Mouse; ss; antisense; complement component C3; inflammation; septic shock; multiple organ failure; hyperacute organ failure; autoimmune disorder; CNS inflammation; multiple sclerosis;
                                                                                                                                             ö
                                                                                                                0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
ive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1. .5
/*tea a /*mod base OTHER
/note= "2'-methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    'note = "2'-methoxyethyl nucleotides"
                                                                                     Sequence 20 BP; 4 A; 6 C; 6 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Location/Qualifiers
                                                                                                                                                                           2495 CCTGCTCAGGCAGCTGG 2514
                                                                                                                                                                                                     20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         /*tag= c
/mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          mod base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Claim 3; Page 27; 72pp; English.
                                                                                                                                                                                            BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      23-OCT-2001; 2001US-00001076.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           23-OCT-2001; 2001US-00001076.
                                                                                                                                                                                                                                                                              ADB90006 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                      (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Ω
                                                                                                                                              Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        6. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                         atherosclerosis; tumour
                                                          developmental disorder.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              . .20
*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 disorder or infection.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2003-606441/57.
                                                                                                                                 Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  US2003096775-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Mus musculus
                                                                                                                                                                                                                                                                                                                                      04-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Graham MJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               22-MAY-2003
                                                                                                                                                                                                                                                                                                         ADB90006;
                                                                                                                   Query Match
   8888888
                                                                                                                                                                                                     g
                                                                                                                                                                                                                                                                                          ð
```

```
complement component C3 and inhibits the expression of complement component C3, or specifically hybridises with at least an 8-nucleobase cortion of an active site on a nucleic acid molecule encoding complement component C3. Also included are a composition comprising complement component C3 in calls or tissues (comprising contacting the component C3 in calls or tissues (comprising contacting the component C3 in calls or tissues (component component C3 in condition associated with complement component C3 comprising administering to the animal the compound cited above so that expression of complement component C3 is inhibited. The antisense compounds are useful for inhibiting the expression of complement C3 is component C3 in cells or tissues, or for treating an animal having a disease or condition associated with complement component C3 such as an expression of complement C3 in cells or tissues, or for treating an animal having a disease or condition associated with complement component C3 such as an experience or autoimmune disorder (e.g. multiple sclerosis), an infection, or attended the complement C3 in cells are also useful as research reagents and diagnostics, in distinguishing functions of various members of a biological pathway, or for preventing or delaying the component sequence is an expension or infection, inflammation. The present sequence is an
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Detector for identifying human papilloma virus subtypes, comprises carrier having two parts carrying first and second oligonucleotides that respectively hybridize with DNA contained in first and second subtypes of the virus.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      The invention comprises oligonucleotides for detecting and identifying subtypes of human papilloma virus (HPV) contained in a sample. The oligonucleotides of the invention are useful for simultaneously detecting and identifying subtypes of HPVs. The present DNA sequence represents an HPV detection oligonucleotide of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human papillomavirus type 33 (HPV 33) detection oligonucleotide #5.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     probe; human papilloma virus; HPV; detection; identification; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Lin Y, Fan C;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Score 15.2; DB 1; Length 20; Pred. No. 2.3e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 5 A; 6 C; 4 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      н, с
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Huang H, Lee B, Lee
Kao Y, Pan C, Chan
                                                                                                                                                                                                                                                                                                                                                                                                             antisense oligonucleotide targeting mouse C3.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Claim 4; SEQ ID NO 131; 221pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1399 TTACCATGAGTTCAAACTTC 1418
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (KING-) KING CAR FOOD IND CO LTD.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1 reaccercacercaaecric 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     10-OCT-2001; 2001EP-00123379.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           10-OCT-2001; 2001EP-00123379
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human papillomavirus type 33
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        17;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 262
        $$$$$$$$$$$$$$$$$$$$$$$$$$$
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      유
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   8
```

Gaps

```
;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            subtypes of human papilloma virus (HPV) contained in a sample. The oligonucleotides of the invention are useful for simultaneously detecting and identifying subtypes of HPVs. The present DNA sequence represents an
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Detector for identifying human papilloma virus subtypes, comprises carrier having two parts carrying first and second oligonucleotides that respectively hybridize with DNA contained in first and second subtypes of
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The invention comprises oligonucleotides for detecting and identifying
                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                    Human papillomavirus type 52 (HPV 52) detection oligonucleotide #6.
                                                                                                                                                                                                                                                                                              probe; human papilloma virus; HPV; detection; identification; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Fan C;
                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   .;
0
                                    0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; tive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
iive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Lin Y,
           Sequence 20 BP; 12 A; 1 C; 2 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence 20 BP; 12 A; 2 C; 2 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Lee H,
Chan P;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     HPV detection oligonucleotide of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Lee B,
Pan C,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Claim 4; SEQ ID NO 263; 221pp; English.
                                                                                         3304 ATTITITITITITITIC 3323
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              3304 ATTTTTTTTTTTTTTTCC 3323
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Huang H,
                                                                                                                                                                                                                                                                                                                                                                                                                                                         (KING-) KING CAR FOOD IND CO LID
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Kao Y,
                                                                                                        20 ATTTTCATTTTATATGTAC 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20 ATTTTCATTTTATATGTGC 1
                                                                                                                                                                                      BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          BP
                                                                                                                                                                                                                                                                                                                                                                                                       10-OCT-2001; 2001EP-00123379.
                                                                                                                                                                                                                                                                                                                                                                                                                                10-OCT-2001; 2001EP-00123379.
                                                                                                                                                                                                                                                                                                                         Human papillomavirus type 52
                                                                                                                                                                                    ADC84033 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADE27866 standard; DNA; 20
                                                                                                                                                                                                                                           (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (first entry)
                                                                Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  You C, 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2003-432398/41.
                                    Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Lin R, Y
Shih C,
                                                                                                                                                                                                                                                                                                                                                  EP1302550-A1.
                                                                                                                                                                                                                                        01-JAN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               29-JAN-2004
                                                                                                                                                                                                                                                                                                                                                                            16-APR-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             the virus.
                                                                                                                                                                                                               ADC84033;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ADE27866;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match
Best Local 8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Lin C,
Hsu H,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 264
                                                                                                                                                          RESULT 263
                                                                                                                                                                         ADC84033/
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADE27866
X S
                                                                                          δ
                                                                                                                  셤
                                                                                                                                                                                                  셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          SAXAX
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ઠે
                                                                                                                                                                                                                                                                                                                                     ô
```

٠<u>٠</u>٠ د ر

```
The invention relates to a method of treating an inflammatory skin disorder in an individual by topically applying an antisense compound targeted to a nucleic acid molecule encoding a human B7 protein. The invention is for treating an inflammatory skin disorder in individual. The skin disorder is psoriasis, contact dermatitis, atopic dermatitis, seborrheic dermatitis, nummular dermatitis, generalised exfoliative dermatitis or czema. The invention effectively modulates critical costimulatory molecules such as the B7 protein. The present sequence represents a human B7-1 targeted oligonucleotide.
                                                                                                                                                                                                                                                                                                                                                        Treating an inflammatory skin disorder such as psoriasis comprises topically applying an antisense compound targeted to the nucleic acid encoding human B7 protein.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         detection; human papillomavirus; HPV subtype; probe; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 20 BP; 7 A; 4 C; 8 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Pred. No. 2.3e
0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                               Example 19; SEQ ID NO 253; 88pp; English.
                                                                                                                                                                                                                                                                                     Karras JG;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  3767 GGCTGGGATCCCTCCCCTGT 3786
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (KING-) KING CAR FOOD IND CO LTD
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20 GGCTGGCATCCCTCTCCTTT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADF43774 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            04-MAY-2001; 2001TW-00110785.
                                                                             2001US-00851871,
                                                                                                                   96US-00777266.
99US-00326186.
                                                                                                                                          04-JUN-1999; 99US-00326186
25-MAY-2000; 2000WO-US014471
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 HPV 33 detecting probe M3305.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Human papillomavirus type 33
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    28-NOV-2001; 2001JP-00362595
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Matches 17; Conservative
                                                                                                                                                                                                                                                                                 Bennett CF, Vickers TA,
                                                                                                                                                                                                   BENNETT C F. VICKERS T A.
                                                                                                                                                                                                                                                                                                                         WPI; 2003-863863/80.
                                                                                                                                                                                                                                           KARRAS J G.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Best Local Similarity
US2003176374-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       JP2002360271-A
                                                                             09-MAY-2001;
                                                                                                                       31-DEC-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           12-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               17-DEC-2002
                                       18-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADF43774;
                                                                                                                                                                                                   (BENN/) H
(VICK/) V
(KARR/) H
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADF43774/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT
ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      The invention relates to a method of treating an inflammatory skin disorder in an individual by topically applying an antisense compound targeted to a nucleic acid molecule encoding a human B7 protein. The invention is for treating an inflammatory skin disorder in individual. The skin disorder is psoriasis, contact dermatitis, atopic dermatitis, seborrabic dermatitis, nummular dermatitis, generalised exfoliative dermatitis or eczema. The invention effectively modulates critical costimulatory molecules such as the B7 protein. The present sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Treating an inflammatory skin disorder such as psoriasis comprises topically applying an antisense compound targeted to the nucleic acid encoding human B7 protein.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                   98; human; B7-1; inflammatory skin disorder; antisense; psoriasis;
contact dermatitis; atopic dermatitis; seborrheic dermatitis;
nummular dermatitis; generalised exfoliative dermatitis; eczema;
critical costimulatory molecule.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ss; human; B7-1; inflammatory skin disorder; antisense; psoriasis; context dermatitis; atopic dermatitis; seborrhaic dermatitis; numular dermatitis; generalised exfoliative dermatitis; eczema; critical costimulatory molecule.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Seguence 20 BP; 3 A; 4 C; 4 G; 9 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.4%; Score 15.2; DB 1;
85.0%; Pred. No. 2.3e+02;
vative 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          represents a human B7-1 targeted oligonucleotide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Human B7-1 targeted oligonucleotide SEQ ID 253.
Human B7-1 targeted oligonucleotide SEQ ID 128
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Example 12; SEQ ID NO 128; 88pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Karras JG;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      TIGAGITICAAAGGITGCIG 1336
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          rrragrircacagcrigcig 20
                                                                                                                                                                                                                                                                                   09-MAY-2001; 2001US-00851871.
                                                                                                                                                                                                                                                                                                                       31-DEC-1996; 96US-00777266.
04-JUN-1999; 99US-00326186.
25-MAY-2000; 2000WO-US014471.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ADE27991 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               29-JAN-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Bennett CF, Vickers TA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                     (VICK/) VICKERS T A.
(KARR/) KARRAS J G.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2003-863863/80.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Query Match
Best Local Similarity
                                                                                                                                                                                                     US2003176374-A1.
                                                                                                                                        Synthetic.
Homo sapiens.
                                                                                                                                                                                                                                             18-SEP-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1317
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ADE27991;
```

SON COURT OF THE STATE OF THE S

ö

Gaps

ö

3; Indels

A detecting apparatus and a detecting method for identifying the subtypes of many species of human papilloma viruses at the same time and a

WPI; 2003-600935/57

Homo sapiens

Synthetic.

RESULT 265

g

Matches

```
ö
                                                                                                                                         This invention describes a novel detecting apparatus for identifying the subtypes of human papillomaviruses (HPV) contained in a sample which comprises a carrier which can load sample, a first oligonucleotide loaded on first part of the carrier and a second oligonucleotide loaded on second part of carrier, in which first and second oligonucleotides hybridise with the DNA of the first and the second HPV subtype and can identify HPV subtype contained in sample at the same time. ADP43644-ADP44289 represent oligonucleotide probes used in the method of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   A detecting apparatus and a detecting method for identifying the subtypes of many species of human papilloma viruses at the same time and a composition for the detection.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       subtypes of human papillomaviruses (HPV) contained in a sample which comprises a carrier which can load sample, a first oligonuclectide loaded on first part of the carrier and a second oligonuclectide loaded on second part of carrier, in which first and second oligonuclectides on the second part of carrier, in which first and the second have subtype and can identify HPV subtype contained in sample at the same time. ADP43644-ADP44289 represent oligonuclectide probes used in the method of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        This invention describes a novel detecting apparatus for identifying the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 2.3e+02; Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        detection; human papillomavirus; HPV subtype; probe; ss
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 12 A; 2 C; 2 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence 20 BP; 12 A; 1 C; 2 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Claim 1; SEQ ID NO 292; 166pp; Japanese.
                                                                      Claim 1; SEQ ID NO 131; 166pp; Japanese.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                3304 ATTTTATTTTATATATCC 3323
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (KING-) KING CAR FOOD IND CO LTD.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20 ATTTCATTTTATATGTAC 1
composition for the detection.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ADF43935 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 HPV 52 detecting probe M5206.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            28-NOV-2001; 2001JP-00362595.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human papillomavirus type 52
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       04-MAY-2001; 2001TW-00110785
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WPI; 2003-600935/57.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              JP2002360271-A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          12-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         17-DEC-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                             invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ADF43935;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 267
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ADF43935/(
11D ADF4;
AC ADF4;
XX ADF4;

셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ð
```

```
The invention relates to a novel polynucleotide isolated and purified from a human gene having any one of 935 fully defined sequences as given in specification, or a sequence having a base substitution. The invention further relates to: an oligonucleotide containing single nucleotide polymorphisms; a PCR primer set chosen from the combination of two DNA fragments from any one of 1220 fully defined sequences as given in specification; a labelling probe containing the SNP containing oligo; and a microarray equipped with the SNP containing oligo. The isolated human gene of the invention is useful for detecting the single nucleotide.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Novel polynucleotide useful for detecting single nucleotide polymorphisms
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   polymorphisms in human gene. The isolated human gene is also useful for diagnosis of disease and determination of side effect to a medical agent. The isolated human gene is also effective in detecting single nucleotide polymorphisms in a human gene. This polymucleotide sequence represents one of the PCR primers used in the single nucleotide polymorphism detection method of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Gapa
                                                                                                                                                                                                                                                     human; single nucleotide polymorphism; microarray; side effect; ss;
primer; PCR.
                                                                                                                                                                                                                          Single nucleotide polymorphism detection primer, SEQ ID No 1648
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
tive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Seguence 20 BP; 5 A; 9 C; 2 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Claim 2; SEQ ID NO 1648; 704pp; Japanese
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (KAGA-) KAGAKU GIJUTSU SHINKO JIGYODAN
3304 ATTTTTTTTTTATATCC 3323
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     2774 CAGGCCTTATGCTAAGGGTG 2793
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20 casscrrrrarsssaassers 1
                     20 ATTTTCATTTTTATATGTGC
                                                                                                                     BP
                                                                                                                                                                                                                                                                                                                                                                                                                                      12-FEB-2002; 2002JP-00034717.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        12-FEB-2002; 2002JP-00034717.
                                                                                                                   ADF88065 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ABZ94003 standard; DNA; 20
                                                                                                                                                                                         (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Local Similarity 85.0 nes 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 2003-820454/77.
                                                                                                                                                                                                                                                                                                                                                                 JP2003235571-A
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                in human gene
                                                                                                                                                                                                                                                                                                                                 Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                17-0CT-2003
                                                                                                                                                                                        26-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                    26-AUG-2003
                                                                                                                                                                                                                                                                                                               Synthetic.
                                                                                                                                                       ADF88065;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ABZ94003;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 269
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ABZ94003,
                                                                                                                                       SXXXEX
                                셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   셤
 ઠ
```

ö

ö

Gaps

.. 0

Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 2.3e+02; Matches 17; Conservative 0; Mismatches 3; Indels

```
23-APR-2002; 2002WO-US013135.
                                                                               24-APR-2001; 2001US-0286137P.
                                                                                        (EPIG-) EPIGENESIS PHARM INC.
                                                                                                                                                                                                                                                                                                                         ABZ86103 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                           17-OCT-2003 (first entry)
                                                                                                                                                                                                                                                                          Local Similarity 85.0
les 17; Conservative
                                                                                                               WPI; 2003-229219/22
                                                                                                 Li Y,
                                                   WO200285308-A2
                                          Homo sapiens.
                                                            31-OCT-2002.
                                                                                                                                        ubiquinone
                                                                                                       Miller S,
                                                                                                                                                                                                                                                                      Query Match
Best Local S
                                                                                                                                                                                                                                                                                                                                  ABZ86103;
                                                                                                 Nyce JW,
                                                                                                                                                                                                                                                                                                               RESULT 270
ABZ86103/c
ID ABZ8611
XX
AC ABZ8611
XX
XX
XX
XX
XX
XX
                                                                                                                                                                                                                                                                                Matches
ð
```

```
The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation codon, coding region, 5' or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiasthmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a use in antisense gene therapy. The composition may have a use in antisense gene therapy. The composition may have a use in antisense year therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of an entisinflammatory steroid in a subject, for reducing or depleting levels of, or reducing sensitivity to adenosine, reducing levels of adenosine receptor, producing bronchodilation, increasing levels of adenosine. In subject, stissue, or treducing levels of ubiquinone or lung surfactant in a subject, stissue, or required profice or condition. Unug allergies, or a respiratory disease or condition. Note: The sequence data for this patent is not represented in the printed profit of appecification, but was obtained in electronic format directly from WIPO
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or
                                              Human, antisense, lung dysfunction, nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiastimmatic; hypotensive; oytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Katz E, Pabalan J, Aguilar
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
.ive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 20 BP; 10 A; 1 C; 3 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      at ftp.wipo.int/pub/published_pct_sequences
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Claim 15; SEQ ID NO 1345; 872pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 3265 TCGAACATGTTCTATTTTT 3284
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ŝ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Li Y, Sandrasagra A,
Tang L, Shahabuddin
Human oligonucleotide sequence.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           24-APR-2001; 2001US-0286137P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (EPIG-) EPIGENESIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                23-APR-2002; 2002WO-US013135.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ABZ89062 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2003-229219/22.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                  WO200285308-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                17-0CT-2003
                                                                                                                                                                                                                                                                          sapiens,
                                                                                                                                                                                                                                                                                                                                                                                           31-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ubiquinone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Nyce JW, I
Miller S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ABZ89062;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ABZ89062/c

ID ABZ890

XX

AC ABZ890

XX

DT 17-OCT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Best Loca
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 271
                                                                                                                                                                                                                                                                          Homo
      ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            임
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonuclectide antisense to the initiation codon, coding region, 5' or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nuclectides of junctions of genes encoding a polypetide associated with lung and/or nasal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiathmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in antisense gene therapy. The composition may have a cuse in artisense gene therapy. The composition may have a cust in a respiratory effect of an antiinflammatory steading or therapeutic respiratory effect of an antiinflammationy bronchodilation, increasing levels of adenosine creeptor, producing bronchodilation, increasing levels of ubiquinone or lung inflammation, lung allergies, or a respiratory disease or condition. Note: The sequence data for this patent is not represented in the printed specification, but was obtained in electronic format directly from WIPO at fig. at ftp.wipo.int/pub/published_pct_sequences
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or
                                                        Human, antisense; lung dysfunction, nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Katz E, Pabalan J, Aguilar D;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.4%; Score 15.2; DB 1; Length 20;
15.0%; Pred. No. 2.3e+02;
ve 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 20 BP; 4 A; 2 C; 2 G; 12 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Disclosure; SEQ ID NO 9245; 872pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Li Y, Sandrasagra A, Ke
Tang L, Shahabuddin S;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1828 ACATAATGACTTCCAAAAA 1847
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20 AAATAATGACTTCAGAAAA 1
      Human oligonucleotide sequence.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      H
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               85.0%;
```

Ö

ö

Gaps

.

Human, antisense, lung dysfunction, nasal airway dysfunction; antiinflammatory, antiallergic; antiinflammatory, antiallergic; antiinflammatory, antiallergic; antisense gene therapy; respiratory; lung, adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds.

Human oligonucleotide sequence

```
Homo sapiens,
            31-OCT-2002
                           ubiquinone.
                    Miller S,
                   Nyce JW,
```

Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or

Disclosure; SEQ ID NO 6827; 872pp; English.

ubiquinone

Katz E, Pabalan J, Aguilar D;

Li Y, Sandrasagra A, K Tang L, Shahabuddin S;

Miller S,

Nyce JW,

WPI; 2003-229219/22.

23-APR-2002; 2002WO-US013135. 24-APR-2001; 2001US-0286137P. (EPIG-) EPIGENESIS PHARM INC.

WO200285308-A2 Homo sapiens.

31-OCT-2002

The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation coodon, coding region, 5 or 3 end genomic flanking regions, 5 and 3 intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an ratinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antialragic, antiasthmatic, hypotenaive, immunosuppressive, and cytostatic activity. The composition may have a cuse in antisense gene therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of an antiinflammatory steroid in a subject, for reducing or depleting levels of of, or reducing sensitivity to adenosine, reducing levels of adenosine receptor, producing bronchodilation, increasing levels of ubiquinone or lung surfactant in a subject of treating by such control of the subject of subject of the subject of subject of the subject of subject of lung inflammation, lung allergies, or a respiratory disease or condition. Note: The sequence data for this patent is not represented in the printed specification, but was obtained in electronic format directly from WIPO Sequence 20 BP; 5 A; 8 C; 4 G; 3 T; 0 U; 0 Other; at ftp.wipo.int/pub/published\_pct\_sequences Disclosure; SEQ ID NO 4304; 872pp; English.

Gaps ö Score 15.2; DB 1; Length 20; Pred. No. 2.3e+02; 0; Mismatches 3; Indels 0.4%; 17; Conservative Query Match Best Local Similarity Matches 17; Conserv

ö

ABZ91585 standard; DNA; 20 ABZ91585; ABZ91585/c ID ABZ9: XX AC ABZ9: XX DT 17-0C

RESULT 272

셤 ઠે

17-OCT-2003 (first entry)

Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or Human; antisense; lung dysfunction; nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; broncholation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds. Li Y, Sandrasagra A, Katz E, Pabalan J, Aguilar D; Tang L, Shahabuddin S; Human oligonucleotide sequence 23-APR-2002; 2002WO-US013135. 24-APR-2001; 2001US-0286137P. (EPIG-) EPIGENESIS PHARM INC. WPI; 2003-229219/22. WO200285308-A2

The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation codon, coding region, 5' or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or masal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiasthmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a tree in antisense gene therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of antiinflammatory steroid in a subject, for reducing levels of adenosine receptor, producing bronchodilation, increasing levels of adenosine receptor, producing bronchodilation, increasing levels of ubiquinone or lung inflammation, lung allergies, or a respiratory disease or condition, lung inflammation, but was obtained in electronic format directly from WIPO at ftp.wipo.int/pub/published\_pct\_sequences 

ö Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 2.3e+02; Matches 17; Conservative 0; Mismatches 3; Indels Sequence 20 BP; 9 A; 3 C; 2 G; 6 T; 0 U; 0 Other; Query Match Best Local S

ö

Gaps

1173 TTAGTTAACTGTAAATTTGG 1192 rrarraacagracarrigg 1 20 셤 ò

RESULT 273 ABZ91732,

1732/c ABZ91732 standard; DNA; 20 ABZ91732;

SAKKEX

(first entry) 17-0CT-2003

```
The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation codon, coling region, 5' or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an entitlenamatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiasthmatic, hypotensive, communosuppressive, and cytostatic activity. The composition may have a cuse in antisense gene therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of an antiinflammatory steroid in a subject, for reducing levels of adenosine creceptor, producing bronchodilation, increasing levels of adenosine creceptor, producing bronchodilation, increasing levels of ubiquinone or lung surfactant in a subject's tissue, or treating bronchoconstriction, lung allergies, or a respiratory disease or condition. Note: The sequence data for this patent is not represented in the printed specification, but was obtained in electronic format directly from WIPO at fig. at ftp.wipo.int/pub/published_pot_sequences
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or
                                   Human, antisense, lung dysfunction, nasal airway dysfunction, antinflammatory steroid; ubiquinone, antinflammatory; antiallergic; antiasthmatic; hypotensive, immunosuppressive, cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy;
                                                                                                                                                                                                                                                                                                                                                                                                                                                Katz E, Pabalan J, Aguilar D;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 11 A; 0 C; 1 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             0.4%; Score 15.2; DB 1;
85.0%; Pred. No. 2.3e+02;
iive 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Disclosure; SEQ ID NO 6974; 872pp; English.
                                                                                                                                                lung inflammation; respiratory disease; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                Li Y, Sandrasagra A, Ka
Tang L, Shahabuddin S;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    3300 ATATTTTTTTTTTATAT 3319
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20 ATATATTTCATATATAT 1
Human oligonucleotide sequence.
                                                                                                                                                                                                                                                                                                                      23-APR-2002; 2002WO-US013135.
                                                                                                                                                                                                                                                                                                                                                              24-APR-2001; 2001US-0286137P.
                                                                                                                                                                                                                                                                                                                                                                                                        (EPIG-) EPIGENESIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ABZ91733 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match
Best Local Similarity 85.0
Marches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2003-229219/22.
                                                                                                                                                                                                                                    WO200285308-A2
                                                                                                                                                                                            Homo sapiens.
                                                                                                                                                                                                                                                                              31-OCT-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ubiquinone
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Miller S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ABZ91733;
                                                                                                                                                                                                                                                                                                                                                                                                                                                  Nyce JW,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT 274
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ABZ91733/c
ID ABZ91
XX
AC ABZ91
XX
DT 17-OC
  g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ò
```

```
The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation codon, coding region, 5' or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiasthmatic, hypotensive, cue in antisense gene therapy. The composition may have a cue in antisense gene therapy. The composition may have a cue in antisense gene therapy. The composition may have a cue in antisense gene therapy. The composition may have a mininflammatory steroid in a subject, for reducing or receptor or therapeutic respiratory effect of an antisinflammatory steroid in a subject, for reducing or depleting levels of committees of composition are producing bronchodilation, increasing levels of adenosine receptor, producing bronchodilation, increasing levels of ubiquinone or lung surfactant in a subject stissue, or treating bronchodilation, lung allergies, or a respiratory disease or condition. Unug inflammation, but was obtained in electronic format directly from WIPO
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Pharmaceutical composition for treating ailments associated with impaired
                               Human; antisense; lung dysfunction; nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  respiration, has oligo(s) antisense to specific gene(s) or its
corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                  Pabalan J, Aguilar D;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Score 15.2; DB 1; Length 20;
Pred. No. 2.3e+02;
0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 20 BP; 11 A; 0 C; 1 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            at ftp.wipo.int/pub/published_pct_sequences
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Disclosure; SEQ ID NO 6975; 872pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                    Katz E,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   3300 ATAITITITITITITI 3319
                                                                                                                                                                                                                                                                                                                                                                                                                                         ŝ
                                                                                                                                                                                                                                                                                                                                                                                                                    Li Y, Sandrasagra A,
Tang L, Shahabuddin
Human oligonucleotide sequence.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ATATATATATATTTTCATAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.4%;
                                                                                                                                                                                                                                                                                                                                         24-APR-2001; 2001US-0286137P.
                                                                                                                                                                                                                                                                                                  23-APR-2002; 2002WO-US013135.
                                                                                                                                                                                                                                                                                                                                                                               (EPIG-) EPIGENESIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ABZ98040 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         17-OCT-2003 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2003-229219/22.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Query Match
Best Local Similarity
                                                                                                                                                                                                                     WO200285308-A2.
                                                                                                                                                                                Homo sapiens
                                                                                                                                                                                                                                                             31-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                    Nyce JW, I
Miller S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ubiquinone
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ABZ98040;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 275
ABZ98040/c
ID ABZ9804
XX
AC ABZ9804
XX
DT 17-OCT-
XX
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Best Loca
Matches
  à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ö
```

. 0

Gaps

ö

17-OCT-2003 (first entry)

Length 20;

```
The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation codon, coding region, 5' or 3' end genomic flanking regions, 5' and 3' incron-exon junctions, 5' or 2' end genomic flanking regions, 5' and 3' incron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or assal airway dysfunction and a second active agent comprising an antiinflammatory, antiallergic, antiasthmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a use in antisense gene therapy. The composition may have a use in antisense gene therapy. The composition may have a use in antisense year therapeutic respiratory effect of an expiratory steptory, lung or therapeutic respiratory effect of an antishmantory steroid in a subject, for reducing levels of denosine receptor, producing bronchodilation, increasing levels of denosine receptor, producing bronchodilation, increasing levels of ubiquinone or lung suffactant in a subject's tissue, or treating bronchoconstriction, lung allergies, or a respiratory disease or condition. Note: The sequence data for this patent is not represented in the printed specification, but was obtained in electronic format directly from WIPO
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or
                                            Human; antisense; lung dysfunction; nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic; antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 20 BP; 5 A; 6 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Disclosure; SEQ ID NO 13282; 872pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          at ftp.wipo.int/pub/published_pct_sequences
                                                                                                                                                               lung inflammation; respiratory disease; ds.
Human MCP4 oligonucleotide sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Li Y, Sandrasagra A, Ka
Tang L, Shahabuddin S;
                                                                                                                                                                                                                                                                                                                                                       23-APR-2002; 2002WO-US013135.
                                                                                                                                                                                                                                                                                                                                                                                                      24-APR-2001; 2001US-0286137P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                  (EPIG-) EPIGENESIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2003-229219/22
                                                                                                                                                                                                                                                          WO200285308-A2
                                                                                                                                                                                                                Homo sapiens.
                                                                                                                                                                                                                                                                                                         31-OCT-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ubiquinone
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Miller S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Nyce JW.
```

```
Gaps
                      ;
0
  Length 20;
                    3; Indels
Score 15.2; DB 1;
Pred. No. 2.3e+02;
0; Mismatches 3;
  0.4%;
                     17; Conservative
           Local Similarity
  Query Match
                      Matches
```

```
3184 TCTCCTTACAGAGGTTAAAG 3203
                     rcrccrrccacacacacaac 1
                     20
 ઠ
                   셤
```

```
ABZ98571 standard; DNA; 20
                                         ABZ98571;
RESULT 276
AB298571
ID AB29857
XX
AC AB29857
XX
DT 17-OCT-
XX
```

17-OCT-2003 (first entry)

Human, antisense, lung dysfunction; nasal airway dysfunction; antiniflammatory; antiallergic; antiniflammatory; antiallergic; antialthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds. Human ICAM oligonucleotide sequence

WO200285308-A2 Homo sapiens.

31-OCT-2002.

23-APR-2002; 2002WO-US013135.

24-APR-2001; 2001US-0286137P.

EPIG-) EPIGENESIS PHARM INC.

Katz E, Pabalan J, Aguilar D;

Katz E, Pabalan J, Aguilar D; Li Y, Sandrasagra A, K Tang L, Shahabuddin S; Miller S, Nyce JW,

WPI; 2003-229219/22.

Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or ubiquinone.

Disclosure; SEQ ID NO 13813; 872pp; English.

The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antiense to the initiation codon, coding regaion, 5 or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an entiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiasthmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a use in antisense gene therapy. The composition may have a use in antisense gene therapy. The composition may have a use in antisense gene therapy. In composition for treating or preventing a respiratory, lung or malignant disease or condition, also for reducing sensitivity to adenosine, reducing levels of an antiinflammatory steroid in a subject, for reducing levels of adenosine, receptor, producing bronchodilation, increasing levels of ubiquinone or lung surfactant in a subject's tissue, or treating bronchoconstriction, lung inflammation, lung allergies, or a respiratory disease or condition.

Note: The sequence data for this patent is not represented in the printed bronch or treating in the printed or the patent is not represented in the printed or the patent is not represented in the printed or the patent is not represented in the printed or the patent is not represented in the printed or the patent is not represented in the printed or the patent is not represented in the printed or the patent is not represented in the printed or the patent is not represented in the printed or the patent is not represented in the printed or the patent is not the patent in the patent in the patent is not the patent in the patent in the patent is not the patent in a for this patent is not represented in the printed obtained in electronic format directly from WIPO at ftp.wipo.int/pub/published\_pct\_sequences 

Gaps ô 0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; iive 0; Mismatches 3; Indels Sequence 20 BP; 5 A; 3 C; 8 G; 4 T; 0 U; 0 Other; 17; Conservative Similarity Query Match Local Matches

ö

1 TTGAGGGGACACAGATGTC 20 AAD52229/c ID AAD52229 standard; DNA; 20 RESULT 277 BXXXEX 요

3750 TTTAGGGAGACACAGATGGC 3769

ઠે

ö

(first entry) 02-MAY-2003

AAD52229;

RESULT 278

```
The compositions comprise antisense compounds, Dempositions and mentions of modulating the expression of interferon gamma receptor 1 (The Compositions comprise antisense compounds, particularly antisense oligonucleotides, tragetted to nucleic acids encoding FRORI. The associated with IPNGRI, such as an autoimmune disorder (e.g. diabetes, autoimmune hypotolis, multiple sclerosis, autoimmune arthritis, autoimmune insulinitis or Crohn's disease), cancer or a disease or condition caused by aberrant apoptosis. It is also used for inhibiting the expression of IFNGRI, as research reagents and diagnostics, to distinguish between functions of various members of a biological pathway, as prophylactic agents (e.g. to prevent or delay infection, inflammation or tumour formation), and as probes or primers. It is also used in the targetted to human IFNGRI DNA. This sequence is used in the exemplification of the invention
                                                                                                                                                                                                                                                 /mod_base= OTHER
/note= "Phosphorothioate backbone; All cytidine residues
                           Human; interferon gamma receptor 1; IFNGR1; autoimmune disorder; cancer; diabetes; autoimmune thyroiditis; multiple sclerosis; immunosuppressive; inflammation; cytostatic; antisense therapy; autoimmune arthritis; autoimmune insulinitis; Crohn's disease; tumour; receptor; antisense; phosphorothioate backbone; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  New antisense oligonucleotides targeted to a nucleic acid molecule encoding interferon gamma receptor 1, useful for treating an autoimmune disorder, e.g. diabetes, multiple sclerosis or Crohn's disease, or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              The invention relates to antisense compounds, compositions and methods
Human IFNGR1 antisense oligonucleotide, ISIS 147645.
                                                                                                                                                                                                                                                                                                                    /*tag= b
/mod base= OTHER
/mote= "2'methoxyethyl nucleotides"
16. .20
                                                                                                                                                                                                                                                                                                                                                                                                                              'note= "2'methoxyethyl nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 20 BP; 4 A; 5 C; 7 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                   are 5-methylcytidines'
                                                                                                                                                                                               Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Claim 3; Page 85; 124pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                              mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        16-APR-2002; 2002WO-US012006.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            26-APR-2001; 2001US-00843376.
                                                                                                                                                                                                                                                                                                                                                                                               υ
                                                                                                                                                                                                                                  *tag=
                                                                                                                                                                                                                                                                                                                                                                                               /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (ISIS-) ISIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Bennett FC, Watt AT;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2003-156687/15.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WO200288162-A1
                                                                                                                                                                                               Key
modified_base
                                                                                                                                                                                                                                                                                                       modified base
                                                                                                                                                                                                                                                                                                                                                                          modified base
                                                                                                                                         Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     07-NOV-2002
                                                                                                                                                              Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           cancer.
```

```
The invention relates to methods for identifying regions of neoplastic growth in a human patient, especially for detecting or predicting metastasis. The methods involve determining whether a neoplastic growth marker protein is overexpressed, either by the use of an antibody consocial protein, or by the use of FCR or hybridisation to detect mucleic acids encoding the marker proteins. A set of neoplastic growth marker: The neoplastic growth marker: The neoplastic growth marker: The neoplastic growth markers are specifically expressed at a higher level in metastatic growth markers are specifically expressed at a higher level in metastatic growth markers are specifically expressed at a higher level in metastatic growth markers with advanced and early stage cancers and normal cells from which the cancer is derived.

CC Overexpression of the neoplastic growth markers is taken as an indication that the tissue has a propensity to metastasise. The invention also encompasses methods for treating aptient with an advanced or metastatic cancer. and for identifying candidate drugs for treating advanced or metastasis, or identifying candidate drugs for treating advanced or detastasis, or identifying candidate drugs for treating advanced or concert in the invention is particularly applicable to destation the neoplastic growth marker proteins are additionally concerted the neoplastic growth marker proteins are additionally concerned the present sequence of the neoplastic growth marker proteins are additionally concerned the present sequence or the present sequence represents a personner measurement and personner or the present sequence represents and the present sequence represents and sequence or the present sequence or the present sequence or 
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      sequence represents a PCR primer used
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Identifying regions of neoplastic growth in a human body, useful for detecting or predicting metastasis, comprises administering to the human body an antibody or peptide that specifically binds to a protein marker of neoplastic growth.
                                                                                                                                                                                                                      Metastasis; neoplastic growth; detection; prediction; neoplastic growth marker; drug screening; cancer; tumour; gastrointestinal; prostate; breast; colorectal; diagnostic imaging; drug targeting; chromosome 8q24.3; human; protein tyrosine phosphatase type IVA member 3; PRL-1; gene mapping;
                                                                                                                                                                        Suman PRL-3 reverse PCR primer #178, used in gene mapping.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 7 A; 6 C; 5 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Bardelli A;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Saha S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         present
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    to map the PRL-3 gene to chromosome
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Disclosure; Page 23; 42pp; English.
                     ADA26894 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    02-OCT-2002; 2002WO-US031247.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        09-OCT-2001; 2001US-0327332P.
                                                                                                                                                                                                                                                                                                                                                                cytostatic; PCR; primer; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ic drugs. The
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Voqelstein B, Kinzler KW,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           SNIXAOH SUHOC VINU ( OLYU)
                                                                                                                       (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 2003-393457/37.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WO2003031930-A2
                                                                                                                                                                                                                                                                                                                                                                                                                 Homo sapiens.
                                                                                                                          20-NOV-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      17-APR-2003.
                                                                      ADA26894;
ADA26894/
```

; 0

Gaps

;

ch 0.4%; Score 15.2; DB 1; Length 20; I Similarity 85.0%; Pred. No. 2.3e+02; 17; Conservative 0; Mismatches 3; Indels

Query Match Best Local Similarity Matches 17; Conserv

ö

Gaps .. 0

Length 20; 3; Indels

Score 15.2; DB 1; Pred. No. 2.3e+02; 0; Mismatches

0.4%;

1689 GAGCAGTTGTCTCCAACAGC 1708

Conservative

Local Similarity es 17; Conserv

Matches

Query Match

20 dagccerrereredadae 1

g ò

```
2071 TGTCGGTCCTCAGTGTGCTT 2090
                       20 rérererereresegaserr
ሯ
                   g
```

ADL16109 standard; DNA; 20 ADL16109; ADL16109

RESULT 279

06-MAY-2004 (first entry)

Human lipoprotein lipase reverse PCR primer R0, SEQ ID NO:6

Human; lipoprotein lipase; LPL; polymorphism analysis; single nucleotide polymorphism; SNP; diagnosis; genetic predisposition; obesity; PCR; primer; ss.

Homo sapiens

KR2002078511-A.

19-OCT-2002,

03-APR-2001; 2001KR-00017640.

03-APR-2001; 2001KR-00017640.

CO LTD. (SEOU-) SEOULIN BIO SCI Lee JJ; Kang BY, Kim GT, Jun GA,

WPI; 2003-338952/32

Primers useful for predicting the likelihood of obesity

Claim 1; SEQ ID NO 6; 7pp; Korean.

The invention relates to PCR primers (ADL16104-ADL16111) for use in the analysis of single nucleotide polymorphisms (SNPs) in the human lipoprotein lipase (PLP) gene (GenBank accession numbers AFG50163) which are associated with a predisposition to obesity. The invention also encompasses a kit comprising a pair of sense and antisense primers. The primers and kit can be used to predict the likelihood of an individual developing obesity, so that it can be prevented. The present sequence represents a specifically claimed human LPL reverse PCR primer of the invention. 

Sequence 20 BP; 4 A; 6 C; 2 G; 8 T; 0 U; 0 Other;

Gaps ô Length 20; 3; Indels Match 0.4%; Score 15.2; DB 1; Local Similarity 85.0%; Pred. No. 2.3e+02; 0; Mismatches 17; Conservative Query Match Matches

ô

2933 GAAAGTCATTTCAACTCTTA 2952 GACAGICITITICACCICITA 20

ð g

AA102454-derived oligonucleotide SEQ ID 6827.

Human, antisense; bronchoconstriction; allergy; hyposecretion; pain; respiratory tract inflammation; adenosine sensitivity; lung; cancer; 

analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory districtses syndrome; albragic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer. surfactant depletion; antiallergic; antiinflammatory; antiasthmatic;

Homo sapiens.

WO200285309-A2

31-OCT-2002.

23-APR-2002; 2002WO-US013143.

24-APR-2001; 2001US-0286036P.

(EPIG-) EPIGENESIS PHARM INC.

Katz E, Pabalan J, Aguilar D; Li Y, Sandrasagra A, Ka Tang L, Shahabuddin S; Nyce JW, I Miller S,

WPI; 2003-093058/08.

S Pharmaceutical composition for treating asthma, has antisense oligonuclectide containing less percentage of adenosine, targeted trucleic acids associated with lung airway or lung dysfunction, and bronchodilating agent.

Claim 15; SEQ ID NO 6827; 763pp; English.

This invention describes a novel composition (a) a first active agent, comprising oligonuclectides, effective for alleviating comprising oligonuclectides, effective for alleviating construction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The coligonuclectides are derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung dysfunction or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonuclectides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention also antiallergic, antiinflammatory, antiasthmatic, analgesic, hypotensive, immunosuppressive and cytostatic activity, is a beta-adrenergic agonist. The composition is useful for preventing or treating a respiratory, ung or malignant disease. The administered composition comprises oligo and is administered to reduce the amount of target polypeptide present in the lungs. The composition or vasilability, or to increase the degradation of the target many or to increase the degradation of the target many or inflammation, allergies and/or bruckctont hypoproduction are associated inflammation, allergies, asthma, impeded respiration, respiratory disease or condition such as pulmonary disease, pulmonary continual disease or condition such as pulmonary disease, pulmonary continual disease or condition such as pulmonary disease, pulmonary continual disease or condition such as pulmonary disease, pulmonary continual disease or condition such as pulmonary disease, pulmonary continual decrease in the target RNA serves to prevent the brackener.

The reduced adenosine content of the anti-sense oligos corresponding to thy information and propertions pulmonary infections, pulmonary entry of prevent the barders and content of th prevent any unwanted effects due to it the system the oligonucleotides into products that free adenosine into 

Sequence 20 BP; 9 A; 3 C; 2 G; 6 T; 0 U; 0 Other;

ö 0.4%; "Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; .ive 0; Mismatches 3; Indels Best Local Similarity 85.0 Matches 17; Conservative

ö

Gaps

8

```
analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer.
                                                                                                                                                                                                                                                                                                    Human, antisense, bronchoconstriction, allergy, hyposecretion, pain, respiratory tract inflammation, adenosine sensitivity, lung, cancer, surfactant depletion, antiallergic, antiinflammatory, antiasthmatic,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           oligonucleotide containing less percentage of adenosine, targeted t
nucleic acids associated with lung airway or lung dysfunction, and
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Katz E, Pabalan J, Aguilar D;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Pharmaceutical composition for treating asthma, has antisense
                                                                                                                                                                                                                                           cathepsin C-derived oligo SEQ ID 1345.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ŝ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sandrasagra A,
                                                              ABD22333 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     23-APR-2002; 2002WO-US013143.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          24-APR-2001; 2001US-0286036P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (EPIG-) EPIGENESIS PHARM INC
                                                                                                                                                                                   29-JUL-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             bronchodilating agent.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2003-093058/08.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Li Y, Sar
Tang L,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WO200285309-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      31-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Miller S,
                                                                                                                          ABD22333;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 3
                                                                                                                                                                                                                                                 Human
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Ношо
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Nyce
RESULT 2

ABD 2333

ABD 2333

ABD 2333

ABD 2333

ABD 2333

ABD 2333

ABD 29-

BB 29-
```

```
This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating comprising oligonucleotides, effective for alleviating bronchoconstriction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung dysfunction or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery of the invention for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antiinflammatory, antiasthmatic, analgesic, hypotensive, immunosuppressive and cytostatic activity, is a beta-adrenergic agonist. The composition is useful for preventing or treating a respiratory, lung or malignant disease. The administered composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypeptide present in the lungs. The composition and/or lung and all administered to reduce the amount of target polypeptide present in the lungs.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             with a disease or condition such as unanary uspontouccion are associated with a disease or condition such as unanary vasoconstriction, inflammation, allergies, asthma, impeded respiration, respiratory distress syndrome, pain, cystic fibrosis, allergic rhinitis, pulmonary transplantation repetition, pulmonary infections, bronchitis, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of the oligonuclectides into products that free adenosine into the system e.g., lung, brain, heart, kidney, etc, tissue environment and thereby, to prevent any unwanted effects due to it
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       inflammation, allergies and/or surfactant hypoproduction are associated
Claim 15; SEQ ID NO 1345; 763pp; English.
```

composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypeptide present in the lungs. The

```
expression of a target polypeptide associated with lung airway or lung dysfunction or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery device, in separate containers; (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antiinflammatory, antiasthmatic, beta-adrenergic agonist. The composition is useful for preventing or treating a respiratory, ing or malignant disease. The administered composition or availability, or to increase the degradation of the target mRNA or to
                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                             Human, antisense; bronchoconstriction, allergy; hyposecretion; pain; respiratory tract inflammation; adenosine sensitivity; lung; cancer; surfactant depletion; antiallergic, antiinflammatory; antiasthmatic; analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; emphysema; chronic obstructive pulmonary disease; pulmonary hypertension; pulmonary transplantation rejection; ss; primer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating bronchoconstriction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene, encoding or regulating
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Pharmaceutical composition for treating asthma, has antisense oligonuclectide containing less percentage of adenosine, targeted to nucleic acids associated with lung airway or lung dysfunction, and
                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Pabalan J, Aguilar D;
                                                                          ö
                                 Score 15.2; DB 1; Length 20; Pred. No. 2.3e+02;
                                                                        3; Indels
Sequence 20 BP; 10 A; 1 C; 3 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                             Human MCP4-derived oligonucleotide SEQ ID 13282
                                                                          0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Claim 15; SEQ ID NO 13282; 763pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Katz E,
                                                                                                                3265 TCGAACATGTTCTATTTTT 3284
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Li Y, Sandrasagra A, K
Tang L, Shahabuddin S;
                                                                                                                                                     -
                                                                                                                                         20 TCAAACAIGCTATATITIT
                                                                                                                                                                                                                                                 ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        23-APR-2002; 2002WO-US013143.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           24-APR-2001; 2001US-0286036P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (EPIG-) EPIGENESIS PHARM INC.
                                       0.4%;
                                                                                                                                                                                                                                   .071/c
ABD31071 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                            (first entry)
                                       Query Match 0.4
Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           bronchodilating agent
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 2003-093058/08.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WO200285309-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens.
                                                                                                                                                                                                                                                                                                                          29-JUL-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  31-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Miller S,
                                                                                                                                                                                                                                                                                         ABD31071;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Nyce JW,
                                                                                                                                                                                                              RESULT 282
                                                                                                                                                                                                                                   ABD31071/
                                                                                                                                                                                                                                                                     g
```

ဌ

dysfunction or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallargic, antiinflammatory, antiasthmatic, antiagesic, hypotensive, immunosuppressive and cytostatic activity, is a beta-adrenergic agonist. The composition is useful for preventing or treating a respiratory, lung or malignant disease. The administered composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or reduce the amount of target polypeptide present in the lungs. The pullmonary obstruction, and/or bronchoconstriction and/or lung with a disease or condition such as pulmonary vasoconstriction.

ô

expression of a target polypeptide associated with lung airway or lung

inflammation, allergies, asthma, impeded respiration, respiratory distress syndrome, pain, cystic fibrosis, allergic rhinitis, pulmonary hypertension, emphysema, chronic obstructive pulmonary disease, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of the oligonucleotides into products that free adenosine into the system

e.g., lung, brain, heart, kidney, etc, tissue environment and thereby, to prevent any unwanted effects due to it

Sequence 20 BP; 5 A; 3 C; 8 G; 4 T; 0 U; 0 Other;

```
inflammation, allergies and/or surfactant hypoproduction are associated with a disease or condition such as pulmonary vasoconstriction, inflammation, allergies, asthma, impeded respiration, respiratory distress syndrome, pain, cystic fibrosis, allergic thinitis, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of the oligonucleotides into products that free adenosine into the system e.g., lung, brain, heart, kidney, etc, tissue environment and thereby, to prevent any unwanted effects due to it
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Human; antisense; bronchoconstriction; allergy; hyposecretion; pain; respiratory tract inflammation; adenosine sensitivity; lung; cancer; surfactant depletion; antiallergic; antiinflammatory; antiasthmatic; analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic fibritis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer.
                                                                                                                                                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                                    ö
pulmonary obstruction, and/or bronchoconstriction and/or lung
                                                                                                                                                                                                                                              0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; ive 0; Mismatches 3; Indels
                                                                                                                                                                                                                Sequence 20 BP; 5 A; 6 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human ICAM-derived oligonucleotide SEQ ID 13813.
                                                                                                                                                                                                                                                                                                                    3184 TCTCCTTACAGAGGTTAAAG 3203
                                                                                                                                                                                                                                                                                                                                                   20 rcrccrrccacacaccrcaac 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               23-APR-2002; 2002WO-US013143.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              24-APR-2001; 2001US-0286036P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (EPIG-) EPIGENESIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                       ABD31602 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       29-JUL-2004 (first entry)
                                                                                                                                                                                                                                                               Local Similarity 85.0
nes 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WO200285309-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              31-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ABD31602;
                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                     Best Loca]
Matches
                                                                                                                                                                                                                                                                                                                                                                                                      RESULT 283
                                                                                                                                                                                                                                                                                                                                                                                                                   ò
                                                                                                                                                                                                                                                                                                                                              ď
```

ö

```
Human, antisense, bronchoconstriction, allergy; hyposecretion; pain, respiratory tract inflammation, adenosine sensitivity; lung; cancer; surfactant depletion; antiallergic, antiinflammatory; antiasthmatic; analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory districtses syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer.
                                  Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Aguilar D;
                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Pharmaceutical composition for treating asthma, has antisense
0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
ative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Pabalan J,
                                                                                                                                                                                                                                                                    AA284245-derived oligonucleotide SEQ ID 9245.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       м,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Katz
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Li Y, Sandrasagra A, Ka
Tang L, Shahabuddin S;
                                                                 3750 TTTAGGGAGACACAGATGGC 3769
                                                                                            1 Trgaeeeeacacacarerc 20
                                                                                                                                                                          ABD30233 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          23-APR-2002; 2002WO-US013143.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       24-APR-2001; 2001US-0286036P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (EPIG-) EPIGENESIS PHARM INC.
                                                                                                                                                                                                                                       29-JUL-2004 (first entry)
 Query Match
Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WPI; 2003-093058/08.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WO200285309-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                               Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           31-OCT-2002.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Miller S,
                                                                                                                                                                                                         ABD30233;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Nyce JW,
                                                                                                                                             RESULT 284
                                                                                                                                                             ABD30233/
                                                                                            g
                                                                                                                                                                                         ò
```

This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating bronchoconstriction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene encoding or regulating

Pharmaceutical composition for treating asthma, has antisense oligonuclectide containing less percentage of adenosine, targeted to nucleic acids associated with lung airway or lung dysfunction, and

Claim 15; SEQ ID NO 13813; 763pp; English.

bronchodilating agent

Katz E, Pabalan J, Aguilar D;

Li Y, Sandrasagra A, Ka Tang L, Shahabuddin S;

Miller S,

Nyce JW,

WPI; 2003-093058/08.

10001863-3.sl.rng

Page 140

```
This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating comprising oligonucleotides, effective for alleviating compositiotion, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, curfactant depoletion or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung dysfunction or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of instruction has antiallergic, antiinflammatory, antiasthmatic, analgesic, hypotensive, immunosuppressive and cytoeratic activity, is a composition composition composition composition are respiratory, lung or malignant disease. The administered composition composition composition and cytoerated to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypeptide present in the lungs. The composition, allergies and/or surfactant hypoproduction are associated with a disease or condition, such as pulmonary vascoconstriction, and/or surfactant hypoproduction and or lung distress syndrome, pain, cystic fibrosis, allergic thinitis, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to the many and the target RNA serves to prevent the breakdown of the oligonucleotides into products that free adenosine and encounted the surfacts of the endenosine content of the endenosine such and the endenosine described adenosine such and such and the endenosine described adenosine endenosine described adenosine described adenosine endenosine described adenosine endenosine such as 
      to
oligonuclectide containing less percentage of adenosine, targeted t
nucleic acids associated with lung airway or lung dysfunction, and
                                                                                                                           Claim 15; SEQ ID NO 9245; 763pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          prevent any unwanted effects due to it
                                                          bronchodilating agent
```

Sequence 20 BP; 4 A; 2 C; 2 G; 12 T; 0 U; 0 Other;

```
/ Match 0.4%; Score 15.2; DB 1; Length 20; Local Similarity 85.0%; Pred. No. 2.3e+02; No 2.3e+02; Conservative 0; Mismatches 3; Indels
                                                                                    1828 ACATAATGACTTCCAAAAA 1847
                                                                                                                               20 AAATAATGACTTCAGAAAA 1
  Query Match
                                              Matches
                                                                                                                            셤
                                                                                      ò
```

ö

Gaps

ö

AA497002-derived oligonucleotide SEQ ID 6975. ABD27963 standard; DNA; 20 BP 29-JUL-2004 (first entry) ABD27963; 

ö

Gaps

beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer. Human, antisense, bronchoconstriction, allergy, hyposecretion, pain, respiratory tract inflammation, adenosine sensitivity, lung, cancer, surfactant depletion, antiallergic, antiinflammatory, antiasthmatic, analgesic, hypotensive, immunosuppressive, cytostatic, cystic fibrosis,

Homo sapiens

WO200285309-A2

31-OCT-2002

This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating comprising oligonucleotides, effective for alleviating comprising oligonucleotides, effective for inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, configuration or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung expression or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antiinflammatory, antiasthmatic, analgesic, hypotensive, immunosuppressive and cytostatic activity, is a contraction as a caspiratory, lung or malignant disease. The administered or educe the amount of target polypeptide present in the lungs. The composition and/or bronchoconstriction and/or lung cinflammation, allergies and/or surfactant hypoproduction are associated distress syndrome, pain, cystic fibrosis, allergic rhinitis, pulmonary chremation, replection, pulmonary infections, bronchits or cancer. The reduced adenosine content of the anti-sense oligos or cancer. The reduced adenosine content of the anti-sense oligos or corresponding to thymidines present in the target RNA serves to prevent the breakdown of the ling process of the oligonic content of the anti-sense oligos or cancer. e.g., lung, brain, heart, kidney, etc, tissue environment and thereby, to prevent any unwanted effects due to it Pharmaceutical composition for treating asthma, has antisense oligonucleotide containing less percentage of adenosine, targeted to nucleic acids associated with lung airway or lung dysfunction, and Human, antisense; bronchoconstriction; allergy; hyposecretion; pain; respiratory tract inflammation; adenosine sensitivity; lung; cancer; surfactant depletion; antiallergic; antiinflammatory; antiasthmatic; Pabalan J, Aguilar D; ô 0.4%; Score 15.2; DB 1; Length 20; 55.0%; Pred. No. 2.3e+02; ve 0; Mismatches 3; Indels Sequence 20 BP; 11 A; 0 C; 1 G; 8 T; 0 U; 0 Other; AI092429-derived oligonucleotide SEQ ID 4304. Katz E, Claim 15; SEQ ID NO 6975; 763pp; English. Tang L, Shahabuddin S; 3300 ATATATTTTTATAT 3319 20 ATATATATATTTTCATAT 1 Sandrasagra A, 뮵 23-APR-2002; 2002WO-US013143. 24-APR-2001; 2001US-0286036P. (EPIG-) EPIGENESIS PHARM INC. 82.0%; ABD25292 standard; DNA; 20 (first entry) Local Similarity 85.0 nes 17; Conservative bronchodilating agent. WPI; 2003-093058/08. Li Y, 29-JUL-2004 Miller S, ABD25292; Query Match Nyce JW, RESULT 286 Matches ABD25292/ g ð

bronchodilating agent 31-OCT-2002. Miller S, Nyce JW, 

beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer. analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; Pharmaceutical composition for treating asthma, has antisense oligonuclectide containing less percentage of adenosine, targeted trucleic acids associated with lung airway or lung dysfunction, and Katz E, Pabalan J, Aguilar D; ii Y, Sandrasagra A, Ka Tang L, Shahabuddin S; 23-APR-2002; 2002WO-US013143. 24-APR-2001; 2001US-0286036P. (EPIG-) EPIGENESIS PHARM INC. WPI; 2003-093058/08 Li Y, WO200285309-A2 Homo sapiens.

Claim 15; SEQ ID NO 4304; 763pp; English

This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating comprising oligonucleotides, effective for alleviating comprising oligonucleotides, readiction, allevials allexials and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, cautefactant depletion or hyposecretion, when administered to a mammal. The coligonucleotides are derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung expression or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antisiflammatory, antisathmatic, analgesic, hypotensive, immunosuppressive and cytostatic activity, is a beta-adrenergic agonist. The composition is useful for preventing or composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to composition compute of target polypeptide present in the lungs. The pulmonary obstruction, and/or bronchoconstriction and/or lung inflammation, allergies and/or surfactant hypoproduction are associated inflammation, allergies and/or surfactant hypoproduction are associated inflammation, allergies and/or surfactant hypoproduction are associated inflammation. inflammation, allergies, asthma, impeded respiration, respiratory distress syndrome, pain, cystic fibrosis, allergic rhinitis, pulmonary hypertension, emphysema, chronic obstructive pulmonary disease, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. ung, brain, heart, kidney, etc, tissue environment and thereby, to The reduced adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of the oligonucleotides into products that free adenosine into the system prevent any unwanted effects due to it

Sequence 20 BP; 5 A; 8 C; 4 G; 3 T; 0 U; 0 Other;

Gaps ö 0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; Live 0; Mismatches 3; Indels Best Local Similarity 85.0 Matches 17; Conservative Query Match

ö

ઠે

ABD27962 standard; DNA; 20 BP. ABD27962/c 

(first entry) 29-JUL-2004

AA497002-derived oligonucleotide SEQ ID 6974.

Human, antisense, bronchoconstriction, allergy, hyposecretion, pain, respiratory tract inflammation; adenosine senstivity; lung; cancer; surfactant depletion; anniallergic, antiinflammatory; antisethmatic; analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic fhintis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer.

Homo sapiens.

WO200285309-A2

31-OCT-2002.

23-APR-2002; 2002WO-US013143.

ç

24-APR-2001; 2001US-0286036P.

(EPIG-) EPIGENESIS PHARM INC

Katz E, Pabalan J, Aguilar D; Li Y, Sandrasagra A, K. Tang L, Shahabuddin S; Miller S, Nyce JW,

WPI; 2003-093058/08.

Pharmaceutical composition for treating asthma, has antisense oligonucleotide containing less percentage of adenosine, targeted to nucleic acids associated with lung airway or lung dysfunction, and oronchodilating agent.

Claim 15; SEQ ID NO 6974; 763pp; English.

This invention describes a novel composition (a) a first active agent, comprising oligonucleotides, effective for alleviating bronchoconstruction, respiratory tract inflammation, allergies and reducing adenosine sensitivity, levels of adenosine (A) or (A) receptors, configurate derived from a gene encoding or regulating expression of a target polypeptide associated with lung airway or lung aypenction also describes a kit, that comprises: (a) a delivery device, in separate containers, (b) the oligonucleotides, (c) instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antiinflammatory, antiactmatic, analgesic, hypotensive, immunosuppressive and cytostatic activity, is a beta-adrenergic agonist. The composition is useful for preventing or treating a respiratory, lung or malignant disease. The administered composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypeptide present in the lungs. The pulmonary obstruction, and/or bronchoconstriction and/or lung inflammation, allergies and/or bronchoconstriction and/or lung with a disease or condition such as pulmonary vasoconstriction, continued, path, cystic fibrosis, allergic rhintis, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to the reduced adenosine content of the anti-sense oligos corresponding to the oligonucleotides into products that free adenosine into the system e.g., lung, brain, heart, kidney, ecc, tissue environment and thereby, to prevent any unwanted effects due to it

RESULT 287

7

```
ADG72205;
                                                                                                                                                                                                                                                                                                     RESULT 289
                                                                                                                                                                                                                                                                                                                ADG72205
                                                                                                                                                                                                                                                                                                                                  g
à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              The invention relates to a compound 8-80 nucleobases in length targeted to, and which specifically hybridises with a nucleic acid molecule encoding sterol regulatory element-binding protein-1 (SRBBP-1, also known as sterol regulatory element-binding transcription factor, SRBBP), and inhibits the expression of SRBBP-1, i.e. is an antisense oligonucleotide. Also included are a compound 8-80 nucleobases in length that specifically hybridises with at least an 8-nucleobase portion of an active site on a nucleic acid molecule encoding sterol regulatory element-binding protein-1, a composition comprising the compound and a carrier or diluent,
                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      New compounds, particularly antisense oligonucleotides targeted to a
nucleic acid encoding sterol regulatory element-binding protein-1, useful
for treating diabetes, atherosclerosis or hyperlipidemia.
                                                                                                                                                                                                                         antisense gene therapy;
sterol regulatory element-binding transcription factor; SREBF;
metabolic disorder; diabetes; cardiovascular disorder; atherosclerosis;
                                                                                                                                                                                                                                                                                                                                  /note= "Phosphorothioate linkages. All cytidines are 5-
                                        Gaps
                                                                                                                                                                                                             Sterol regulatory element-binding protein-1; SREBP-1; ss; human;
                                        ;
0
                 Score 15.2; DB 1; Length 20;
Pred. No. 2.3e+02;
0; Mismatches 3; Indels
                                                                                                                                                                                          Human SREBP-1 antisense oligonucleotide ISIS 220068.
                                                                                                                                                                                                                                                                                                                                                                            /mod_base= OTHER
/note= "2'-methoxyethtyl residues"
 Sequence 20 BP; 11 A; 0 C; 1 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                           'note= "2'-methoxyethyl residues"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Example 15; SEQ ID NO 66; 112pp; English.
                                                                                                                                                                                                                                                                                             Location/Qualifiers
                                                            3300 ATATATTTTATTT 3319
                                                                                                                                                                                                                                                                                                                                                                                                         /*tag= c
/mod_base= OTHER
                                                                                                                                                                                                                                                                                                                         base= OTHER
                                                                                                                                                                                                                                                                                                                                                                          base= OTHER
                                                                                                                                                                                                                                                                                                                                              methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Dobie KW;
                                                                        ATATATTTTCATATATAT 1
                      0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       04-JUN-2002; 2002US-00161996
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         04-JUN-2002; 2002US-00161996
                                                                                                                                ADG72071 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                  Д
                                                                                                                                                                                                                                                                                                                                                                   ๙
                                                                                                                                                                        (first entry)
                                         17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                              16. .20
                                                                                                                                                                                                                                                                                                                                                                 /*tag=
                                                                                                                                                                                                                                                                                                        ...20
*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Baker BF,
                                                                                                                                                                                                                                                                                                                           mod/
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 2004-022079/02.
                       Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                US2003224515-A1
                                                                                                                                                                                                                                                      hyperlipidaemia
                                                                                                                                                                                                                                                                                               Key
modified_base
                                                                                                                                                                                                                                                                                                                                                         modified_base
                                                                                                                                                                                                                                                                                                                                                                                               modified base
                                                                                                                                                                        11-MAR-2004
                                                                                                                                                                                                                                                                            Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   04-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Freier SM,
                                                                                                                                                     ADG72071;
                                                                                20
                                         Matches
                                                                                                               RESULT 288
                                                                                                                         ADG72071
                                                                                                                                  à
                                                                                요
```

```
inhibiting the expression of sterol regulatory element-binding protein-1

con cells or tissues (by contacting the cells or tissues with the compound

contact expression of sterol regulatory element-binding protein-1 is

conhibited) and treating an animal having a disease or condition

casociated with sterol regulatory element-binding protein-1 by

compound so that expression of sterol regulatory element-binding protein-1

compound so that expression of sterol regulatory element-binding protein-

compound so that expression of sterol regulatory element-binding protein-

compound so that expression of sterol regulatory element-binding protein-

compound so that expression of sterol regulatory element-binding protein-1, such as a more modified nucleobase (preferably 2.0-methoxyethyl sugar

compety) or at least one modified nucleobase (preferably 5.

methylcytosine). The compound, composition and methods are useful for

treating a disease or condition associated with sterol regulatory element

contacting protein-1, such as a metabolic disorder e.g. diabetes, or a binding protein-1, such as a metabolic disorder e.g. diabetes, or a cardiovascular disorder, e.g. atherosclerosis or hyperlipidaemia. They

are also useful in research and diagnostics for modulating the expression of sterol regulatory element-binding protein-1. The present sequence is
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       The invention relates to a compound 8-80 nucleobases in length targeted to, and which specifically hybridises with a nucleota acid molecule encoding sterol regulatory element-binding protein-1 (SREBP-1, also known as sterol regulatory element-binding transcription factor, SREBF), and inhibits the expression of SREBP-1, i.e. is an antisense oligonucleotide.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              New compounds, particularly antisense oligonucleotides targeted to a nucleic acid encoding sterol regulatory element-binding protein-1, useful for treating diabetes, atherosclerosis or hyperlipidemia.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            julatory element-binding transcription factor; SREBF; disorder; diabetes; cardiovascular disorder; atherosclerosis;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sterol regulatory element-binding protein-1; SREBP-1; ss; human;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      0.4%; Score 15.2; DB 1; Length 20; 35.0%; Pred. No. 2.3e+02; ve 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      an antisense oligonucleotide targeting human SREBP-1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 20 BP; 3 A; 3 C; 7 G; 7 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Example 16; SEQ ID NO 200; 112pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   8 CACTGCTGCTCACAGAAGCA 27
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Dobie KW
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20 cacrecrerceacaaagea 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ВР
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Human SREBP-1 target site #42
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            04-JUN-2002; 2002US-00161996.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          04-JUN-2002; 2002US-00161996
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      85.0%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADG72205 standard; cDNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
Best Local Similarity 85.0°
Marches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              sterol regulatory elementerapy; sterol regulatory elementabolic dierrity elementerapolic dierrity elementerapolic dierrity elementerapolic dierrity elementerapolic dierrity elementerapolic dierrity elementerapolic dierrity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WPI; 2004-022079/02.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          hyperlipidaemia.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              US2003224515-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           11-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                04-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Freier SM,
```

```
hybridises with at least an 8-nucleobases in length that specifically hybridises with at least an 8-nucleobase portion of an active site on a nucleic acid molecule encoding sterol regulatory element-binding protein-1, a composition comprising the compound and a carrier or diluent, in cells or tissues (by contacting the cells or tissues with the compound so that expression of sterol regulatory element-binding protein-1 is contacted with sterol regulatory element-binding protein-1 is ansociated with sterol regulatory element-binding protein-1 by administering to the animal a therapeutic or prophylactic amount of the compound so that expression of sterol regulatory element-binding protein-1 by administering to the animal at therapeutic or prophylactic amount of the compound so that expression of sterol regulatory element-binding protein-1 is inhibited. The antisense oligonuclectide comprises at least one modified sugar moiety (preferably 2'-0-methoxyethyl sugar moiety) or at least one modified nucleobase (preferably 5-methoxyethyl sugar moiety) or at least one modified nucleobase (preferably 5-methoxyethyl sugar creating a disease or condition associated with sterol regulatory element to treating a disease or condition as a metabolic disorder e.g. diabetes, or a cardiovascular disorder, e.g. atherosclerosis or hyperlipidaemia. They are also useful in research and diagnostics for modulating the expression of thuman SREBP-1 target region for the antisense oligonucleotides.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Sequence 20 BP; 7 A; 7 C; 3 G; 3 T; 0 U; 0 Other;
```

0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; 3; Indels 0; Mismatches 8 CACTGCTGCTCACAGAAGCA 27 CACTGCTGTCCACAAAGCA 20 Local Similarity 85.0 nes 17; Conservative Query Match Matches

ઠે 셤

ö

Gaps

ö

Human glucocorticoid receptor-specific antisense oligonucleotide #4377. ADH67543 standard; DNA; 20 BP. ADH67543; RESULT 290 

25-MAR-2004 (first entry)

antisense oligonucleotide; glucocorticoid receptor; infection; inflammation; tumour formation; diabetees; obesity; cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss; phosphorothioate backbone; 2'-methoxyethyl; 2'-MOB.

Homo sapiens

WO2003099215-A2

04-DEC-2003.

20-MAY-2003; 2003WO-US016084,

20-MAY-2002; 2002US-0381857P.

(PHAA ) PHARMACIA CORP

Nalseth AE; Crosby SD,

WPI; 2004-035034/03.

New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, mammalian glucocorticoid receptor, useful for treating diabete cardiovascular disorder, hyperlipidemia or Cushing's syndrome

Claim 4; SEQ ID NO 4377; 985pp; English.

The invention comprises an antisense oligonucleotides that are targeted to nucleic acids encoding a mammalian glucocorticoid receptor. The

3302 ATATTTTATTTTATATAT 3321

ò

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human gluccorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The invention comprises an antisense oligonucleotides that are targeted antisense acids encoding a mammalian glucocorticoid receptor. The antisense oligonucleotides of the invention are useful for preventing or delaying infection, inflammation or tumour formation. The antisense oligonucleotides are also useful for treating diabetes, obesity, cardiovascular disorders, hyperlipidaemia or Cushing's syndrome. The present DNA sequence represents an antisense oligonucleotide that targets the human glucocorticoid receptor gene. NOTE: The present sequence contains 2'-methoxyethyl (2'-MOE) wings and a phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            New antisense compound targeted to a nucleic acid molecule encoding mammalian glucocorticoid receptor, useful for treating diabetes, obesity, cardiovascular disorder, hyperlipidemia or Cushing's syndrome.
    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            antisense oligonucleotide; glucocorticoid receptor; infection;
hifiammation; tumour formation; diabetes; obesity;
cardiovascular disorder; hyperlipidaemia; Cushing's syndrome; human; ss;
phosphorothioate backbone; 2'-methoxyethyl; 2'-MOE.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human glucocorticoid receptor-specific antisense oligonucleotide #4661
antisense oligonucleotides of the invention are useful for preventing
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ô
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                / Match 0.4%; Score 15.2; DB 1; Length 20; Local Similarity 85.0%; Pred. No. 2.3e+02; nes 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                             3; Indels
                                                                                                                                                                                                0.4%; Score 15.2; DB 1; Length
85.0%; Pred. No. 2.3e+02;
tive 0; Mismatches 3; Indels
                                                                                                                                                               Sequence 20 BP; 13 A; 1 C; 0 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 20 BP; 13 A; 1 C; 0 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Claim 4; SEQ ID NO 4661; 985pp; English.
                                                                                                                                                                                                                                                                                   3301 TATATTTTTTTTATA 3320
                                                                                                                                                                                                                                                                                                                            20 rararrrrararrragrra 1
                                                                                                                                                                                                                                                                                                                                                                                                                               BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20-MAY-2003; 2003WO-US016084,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 20-MAY-2002; 2002US-0381857P.
                                                                                                                                                                                                                                                                                                                                                                                                                  827/c
ADH67827 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                25-MAR-2004 (first entry)
                                                                                                                                                                                                     Query Match
Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Crosby SD, Nalseth AE;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2004-035034/03.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WO2003099215-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   04-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ADH67827;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                                                                           RESULT 291
                                                                                                                                                                                                                                                                                                                                                                                                              ADH67827,
  8
                                                                                                                                                                                                                                                                                       ઠે
                                                                                                                                                                                                                                                                                                                       셤
```

g

20

```
The present invention relates to antisense compounds targeted to a nucleic acid encoding interleukin-1 (IL-1) receptor-associated kinase-1 (IRA-1). The antisense compound comprises an antisense oligonucleotide that specifically hybridises with the nucleic acid and inhibits the expression of IRAK-1. The antisense oligonucleotide is a chimeric oligonucleotide. The antisense oligonucleotide comprises at least one addition internucleoside linkage, preferably a phosphorothioate linkage. The also comprises at least one modified sugar moiety, preferably a 2'-0- methoxyethyl (2'-MOB) sugar moiety. The antisense oligonucleotide further comprises at least one modified nucleobase, preferably a 5- methoxyethyl (2'-MOB) sugar moiety. The antisense oligonucleotide further methylcytosine. The antisense oligonucleotides are useful for the reatment of diseases such as hyperproliferative disorders, e.g. cancer, autoimmune disorders, altered bone metabolism, and inflammation. The present sequence represents an antisense oligonucleotide used in the examples of the present invention.
                                                                                                                                                                                                                                                                                                                                                              Antisense therapy; human; interleukin-1 receptor-associated kinase-1; IL-1 receptor-associated kinase-1; IRAK-1; hyperproliferative disorder e.g.; cancer; autoimmune disorder; altered bone metabolism or inflammation; cytostatic; immunosuppressive; osteopathic; antiinflammatory; phosphorothioate; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /mod_base= OTHER //mod_base a phosphorothioate //note= "This oligonuclectide has a phosphorothioate backbone and 2'-methyoxyethyl (2'-MOE) wings at the 5' and 3' ends, which are 5 mucleotides in length at each end. All cytidine residues are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             New compound having a sequence targeted to a nucleic acid encoding II receptor-associated kinase-1, useful for preparing a composition for treating hyperproliferative or autoimmune disorder or inflammation.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 4 A; 6 C; 7 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                               Human IRAK-1 DNA, antisense oligonucleotide #49.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Example 15, SEQ ID NO 62; 66pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              10-JUN-2002; 2002US-00167034
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     10-JUN-2002; 2002US-00167034
                                                                      ADH50655 standard; DNA; 20
                                                                                                                                                                                                                            (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ø
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Freier SM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2004-052028/05.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         JS2003228690-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Key
modified_base
                                                                                                                                                                                                                        25-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    11-DEC-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Baker BF,
                                                                                                                                                 ADH50655;
RESULT 292
                               ADH50655/

ID ADH5

ADH5055/

ADH50655/

ADH5065/

ADH50
```

```
The present invention relates to antisense compounds targeted to a nucleic acid encoding interleukin-1 (IL-1) receptor-associated kinase-1 (IRAK-1). The antisense compound comprises an antisense oligonucleotide that specifically hybridises with the nucleic acid and inhibits the expression of IRAK-1. The antisense oligonucleotide is a chimeric oligonucleotide. The antisense oligonucleotide is a chimeric comprises at least one modified comprises at least one modified sugar moiety, preferably a 2'-0- It also comprises at least one modified sugar moiety, preferably a 2'-0- methoxyethy! (2'-MOE) sugar moiety. The antisense oligonucleotide further comprises at least one modified nucleobase, preferably a 2'-0- comprises at least one modified nucleobase, preferably a 2'-0- comprises at least one modified nucleobase, preferably a 2'-0- comprises at least one modified nucleobase, preferably a 2'-0- comprises at least one modified nucleobase, preferably a 2'-0- comprises at least one modified nucleobase, preferably a 2'-0- comprise at least one modified nucleobase, preferably a 2'-0- comprise at least one modified nucleobase, preferably a 2'-0- comprise at least one modified nucleobase, preferably a 2'-0- comprise at least one metabolism, and inflammation. The present sequence represents a human IRAK-1 DNA target sequence for an
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    New compound having a sequence targeted to a nucleic acid encoding IL-1 receptor-associated kinase-1, useful for preparing a composition for treating hyperproliferative or autoimmune disorder or inflammation.
                                                                                                                                                                                                                                                                                                   Antisense therapy; human; interleukin-1 receptor-associated kinase-1; IL-1 receptor-associated kinase-1; IRAK-1; hyperproliferative disorder e.g.; cancer; autoimmune disorder; altered bone metabolism or inflammation; cytostatic; immunosuppressive; osteopathic; antiinflammatory; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          .;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; tive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Sequence 20 BP; 3 A; 7 C; 6 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Example 15; SEQ ID NO 128; 66pp; English.
                                                                                                                                                                                                                                                                         Human IRAK-1 DNA target sequence #43
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 460 AGAGCCTAAGCCACCTCTCT 479
460 AGAGCCTAAGCCACCTCTCT 479
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AGAGCCTAGGCGGCCTCTCT 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Dobie KW;
                            20 AGAGCCTAGGCGGCCTCTCT 1
                                                                                                                                                 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     10-JUN-2002; 2002US-00167034.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              10-JUN-2002; 2002US-00167034.
                                                                                                                                                 ADH50721 standard; DNA; 20
                                                                                                                                                                                                                                  (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    antisense oligonucleotide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Freier SM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2004-052028/05.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        US2003228690-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                               Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     SISI (-SISI)
                                                                                                                                                                                                                                  25-MAR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             11-DEC-2003.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Baker BF,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             н
                                                                                                                                                                                             ADH50721;
                                                                                                            RESULT 293
                                                                                                                                  ADH50721
                                                                                                                                                    ð
                                     셤
      ઠે
```

ADI27028 standard; DNA; 20 BP.

RESULT 294 ADI27028/c ID ADI270

ö

0; Gaps

Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 2.3e+02; Matches 17; Conservative 0; Mismatches 3; Indels

```
The invention describes a new antisense oligonucleotide, having a sequence comprising 8-80 bp targeted to a nucleic acid encoding cyclindependent kinase 4, specifically hybridises with the nucleic acid encoding cyclin-dependent kinase 4 and inhibits expression of cyclindependent kinase 4. The antisence oligonucleotide is useful for preparing a composition for treating diabetes, infertility or hyperproliferative disorder, e.g., cancer. This sequence represents a human cyclin dependent
                                                                                                                                                                                                                                                                                           /mod_base= OTHER
/note= "OTHER= Phosphorothioate backbone. All cytidines
are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         New antisense oligonucleotide, having a sequence targeted to a nucleic acid encoding cyclin-dependent kinase 4, useful for preparing a composition for treating diabetes, infertility or hyperproliferative
                                                                                                                                                                                                                                                                                                                                                                                            /mod_base= OTHER
/note= "OTHER=.2'-O-Methoxyethyl (2'-MOE) nucleotides'
                                                                                                                                                                                                                                                                                                                                                                                                                                                                          /mod_base= OTHER
/note= "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides'
                                                                                                             cytostatic, antidiabetic; antiinfertility; gene therapy;
cyclin-dependent kinase 4; diabetes; infertility;
hyperproliferative disorder; cancer; antisense technology; human; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
ive 0; Mismatches 3; Indel8
                                                                             Cyclin dependent kinase 4 antisense oligonucleotide #54.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 20 BP; 8 A; 2 C; 6 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Example 15; SEQ ID NO 73; 90pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          kinase 4 antisense oligonucleotide.
                                                                                                                                                                                                                                    Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   3592 GCTATAGGCATGAAGGAAGT 3611
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1 GCAATTGGCATGAAGGAAAT 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       02-JUL-2002; 2002US-00188779.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                02-JUL-2002; 2002US-00188779
                                        (first entry)
                                                                                                                                                                                                                                                                          Ω
                                                                                                                                                                                                                                                                                                                                                                            *tag= a
                                                                                                                                                                                                                                                                                                                                                                                                                                                     /*tag= c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Local Similarity 85.0
nes 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                   15. .20
                                                                                                                                                                                                                                                        ...20
*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         disorder, e.g., cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Dean NM, Freier SM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2004-081710/08
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     US2004005567-A1
                                                                                                                                                                                                                                      Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                 modified_base
                                                                                                                                                                                                                                                                                                                                                        modified base
                                                                                                                                                                                               Homo sapiens
                                        22-APR-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           08-JAN-2004
  ADI26888;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Best Loca
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 296
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ADJ34054/
8,8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    The invention describes a new antisense oligonucleotide, having a sequence comprising 8-80 bp targeted to a nucleic acid encoding cyclindependent kinase 4, specifically hybridises with the nucleic acid encoding cyclin-dependent kinase 4 and inhibits expression of cyclindependent kinase 4. The antisense oligonucleotide is useful for preparing a composition for treating diabetes, infertility or hyperproliferative disorder, e.g., cancer. This sequence represents a human cyclin dependent kinase 4 antisense oligonucleotide.
                                                                                                                                                                                                                                                                                                         /mod_base= OTHER
/note= "OTHER= Phosphorothioate backbone. All cytidines
are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              New antisense oligonucleotide, having a sequence targeted to a nucleic acid encoding cyclin-dependent Kinase 4, useful for preparing a composition for treating diabetes, infertility or hyperproliferative disorder, e.g., cancer.
                                                                                                                                                                                                                                                                                                                                                                                            /*tag= a
/mod base= OTHER
/mod = "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides"
15. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /note= "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides
                                                                                                                                cytostatic; antidiabetic; antiinfertility; gene therapy;
cyclin-dependent kinase 4; diabetes; infertility;
hyperproliferative disorder; cancer; antisense technology; human; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Indels
                                                                                             Cyclin dependent kinase 4 antisense oligonucleotide #194
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 4 A; 6 C; 2 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Example 15; SEQ ID NO 213; 90pp; English.
                                                                                                                                                                                                                                                      Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       3592 GCTATAGGCATGAAGGAAGT 3611
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            J= c
base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            20 GCAATTGGCATGAAGGAAAT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  02-JUL-2002; 2002US-00188779
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         02-JUL-2002; 2002US-00188779
                                                      22-APR-2004 (first entry)
                                                                                                                                                                                                                                                                                             Ω
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Conservative
                                                                                                                                                                                                                                                                          . 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Dean NM, Freier SM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2004-081710/08:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       US2004005567-A1
                                                                                                                                                                                                                                                        Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                          modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                       modified base
                                                                                                                                                                                                                   Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              08-JAN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 17:
                    ADI27028;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     RESULT 295
AD126888
ID AD1268
```

ö

Gaps

ö

ADJ34054 standard; DNA; 20

ADJ34054

ADI26888 standard; DNA; 20 BP.

Matches

ሯ 유 Š

polo-like kinase; polo-like kinase inhibitor; antisense oligonucleotide; cytostatic; antiinflammatory; antimicrobial; antisense gene therapy; knase inhibitor; hyperproliferative disorder; ancer; non-small cell lung cancer; osesphagaal cancer; infection; inflammation; tumour; human; phosphorothioate; 2'-0-methoxyethyl; ss.

Homo sapiens.

Synthetic.

Human polo-like kinase antisense oligonucleotide SEQ ID NO:55.

(first entry)

22-APR-2004

ADJ33995;

BP

ADJ33995 standard; DNA; 20

20 TGGTTCTTTTTTCCCGGG

```
셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      The present invention describes a compound (1) of 8-80 nucleobases in length targeted to a nucleic acid molecule encoding polo-like kinase, where (1) specifically hybridises with nucleic acid molecule encoding polo-like kinase and inhibits the expression of polo-like kinase, or specifically hybridises with at least an 8-nucleobase portion of a preferred target region on a nucleic acid molecule encoding polo-like kinase. Also described: (1) a composition comprising (1) and a planamaceutical carrier or diluent; (2) inhibiting the expression of polo-like kinase in cells or tissues comprising contacting the cells or tissues with (1); (3) treating an animal having a disease or condition associated with polo-like kinase comprising administering to the animal at the rapeutic or prophylactic amount of (1) so that expression of polo-like kinase is inhibited; and (4) screening for an antisense compound comprising a preferred target region of a nucleic acid molecule encoding polo-like kinase with one or more candidate antisense compounds with one or more candidate antisense compounds which inhibits the expression of a compounds comprising at least an 8-nucleobase portion which is complementary to the preferred target region, and selecting for one or more candidate antisense compounds which inhibits the expression of a cutiented antisense compounds which inhibits. The antisense contolides or compounds (1) can be used for inhibiting the artisense decompounds (1) can be used for inhibiting the expression of polo-like kinase, and can be used in a sesociated with aberrant expression of polo-like kinase, including non-small cell ung research reagents and kits, or in diagnostic, therapeutic applications, e.g. to prosphylactic applications, e.g. to prosphylactic applications, e.g. to present or despay infection, inclummation or tumour formation. The present sequence represents a human of polo-like presents and entime or inflammation or tumour formations.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                New antisense compounds targeted to nucleic acid molecules encoding polo-
like kinae, useful for treating diseases associated with aberrant
expression of polo-like kinase, e.g. non-small cell lung cancer or
esophageal cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    polo-like kinase target oligonucleotide, which is used in an example from
                                                                                                                                                                                                         polo-like kinase; polo-like kinase inhibitor; antisense oligonucleotide; cytostatic; antiinflammatory; antimicrobial; antisense gene therapy; kinase inhibitor; hyperproliferative disorder; cancer; non-small cell lung cancer; oesophageal cancer; infection; inflammation; tumour; human; target; se.
                                                                                                    Human polo-like kinase target oligonucleotide SEQ ID NO:114.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 20 BP; 10 A; 5 C; 5 G; 0 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Example 15; SEQ ID NO 114; 138pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  25-JUL-2003; 2003WO-US023413.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            30-JUL-2002; 2002US-00209405.
22-APR-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  the present invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Wyatt JR, Freier SM;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2004-143840/14.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WO2004011610-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             05-FEB-2004.
```

```
The present invention describes a compound (1) of 8-80 nucleobases in length targeted to a nucleic acid molecule encoding polo-like kinase, where (1) specifically hybridises with nucleic acid molecule encoding polo-like kinase and inhibits the expression of polo-like kinase, or specifically hybridises with at least an 8-nucleobase portion of a preferred target region on a nucleic acid molecule encoding polo-like kinase. Also described: (1) a composition comprising (1) and a charmaceutical carrier or diluent; (2) inhibiting the expression of polo-like kinase in cells or tissues comprising contacting the calls or tissues with (1); (3) treating an animal having a disease or condition associated with polo-like kinase comprising administering to the animal a charpettic or prophylactic amount of (1) so that expression of polo-like kinase is inhibited; and (4) screening for an antisense compound comprising contacting a preferred target region of a nucleic acid molecule encoding polo-like kinase with one or more candidate antisense
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            New antisense compounds targeted to nucleic acid molecules encoding polo-
like kinase, useful for treating diseases associated with aberrant
expression of polo-like kinase, e.g. non-small cell lung cancer or
esophageal cancer.
                                                                                                                                                                                                                                                                                                                         note = "phosphorothioate linkages"
                                                                                                                                                                                                                                                                                                                                                                                      /note= "2'-0-methoxyethyls"
16. .20
                                                                                                                                                                                                                                                                                                                                                                                                                                                 /note= "2'-0-methoxyethyls"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Claim 1; SEQ ID NO 55; 138pp; English.
                                                                                                                                                                                                                                                            Location/Qualifiers
                                                                                                                                                                                                                                                                                                         'mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                    /*tag= c
/mod base= OTHER
                                                                                                                                                                                                                                                                                                                                                                      /mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           25-JUL-2003; 2003WO-US023413.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      30-JUL-2002; 2002US-00209405
                                                                                                                                                                                                                                                                                          Ω.
                                                                                                                                                                                                                                                                                                                                                          ಹ
                                                                                                                                                                                                                                                                             ...20
*tag=
                                                                                                                                                                                                                                                                                                                                                       *tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (ISIS-) ISIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Freier SM;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 2004-143840/14.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WO2004011610-A2.
                                                                                                                                                                                                                                                                            modified base
                                                                                                                                                                                                                                                                                                                                         modified base
                                                                                                                                                                                                                                                                                                                                                                                                      modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             05-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Wyatt JR,
```

ó,

Gaps

ö

Score 15.2; DB 1; Length 20; Pred. No. 2.3e+02; 0; Mismatches 3; Indels

0.4%;

Local Similarity 85.0 nes 17; Conservative

Query Match Matches

compounds comprising at least an 8-nucleobase portion which is complementary to the preferred target region, and selecting for one or more candidate antisense compounds which inhibits the expression of a nucleic acid molecule encoding polo-like kinase. (1) has cytostatic, antisense gene therapy, and as a kinase inhibitor. The antisense antisense gene therapy, and as a kinase inhibitor. The antisense conjugance or compounds (1) can be used for inhibiting the expression of polo-like kinase, and for treating diseases or conditions associated with aberrant expression of polo-like kinase, e.g. hyperproliferative disorder such as cancer, including non-small cell lung cancer or oesophageal cancer. The antisense compounds are also useful as research reagents and kits, or in diagnostic, therapeutic and prophylactic applications, e.g. to prevent or delay infection, inflammation or tumour formation. The present sequence represents a human polo-like kinase chimeric phosphorchicate antisense oligonucleotide, which is used in an example from the present invention.

Sequence 20 BP; 0 A; 5 C; 5 G; 10 T; 0 U; 0 Other;

Score 15.2; DB 1; Length 20; Pred. No. 2.3e+02; 0; Mismatches 3; Indels 1091 TGTTTCTTCATTTTCCCTGG 1110 TGGTTCTTCTTTTTCCCGGG 20 Query Match 0.4%; Best Local Similarity 85.0%; Matches 17; Conservative ( ò 셤

ADJ46575 standard; DNA; 20 BP

ADJ46575;

(first entry) 06-MAY-2004 Human requiem antisense oligonucleotide ISIS #204782

human; requiem; hyperproliferative disorder; cancer; developmental disorder; infection; inflammation; tumour formation; ss; antisense.

Homo sapiens.

Synthetic.

05-FEB-2004

US2004023385-A1.

05-AUG-2002; 2002US-00212993

05-AUG-2002; 2002US-00212993

(ISIS-) ISIS PHARM INC

Bennett CF, Freier SM,

Dobie KW

WPI; 2004-142666/14.

New antisense compound targeted to a nucleic acid molecule encoding requiem, useful for modulating expression of requiem or for treating cancer or developmental disorders.

Example 15; SEQ ID NO 50; 66pp; English.

antisense compound can also be used as research tools and diagnostics. It can also be used as tools in differential and/or combinatorial analyses to elucidate expression patterns of a portion or the entire complement of The invention relates to a compound targeted to a nucleic acid molecule encoding requiem which specifically hybridises with the nucleic acid molecule encoding requiem and inhibits the expression of requiem. The compound, particularly the antisense oligonucleotide is useful in modulating the function of nucleic acid molecules encoding requiem. The 

for treating diseases or conditions associated with requiem, preferably hyperproliferative disorder, e.g. cancer or a developmental disorder. The compound can also be used as prophylaxis, e.g. to prevent or delay inflection, inflammation or tumour formation. The present sequence represents the human requiem antisense oligonucleotide. expressed within cells and tissues. The compound can also be used 858888888

Seguence 20 BP; 4 A; 8 C; 4 G; 4 T; 0 U; 0 Other;

Gaps ö 0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; ative 0; Mismatches 3; Indels Local Similarity 85.0 les 17; Conservative Query Match Best Loca Matches

ö

ò 셤 RESULT 299

BP ADJ61423 standard; DNA; 20

ADJ61423;

ô

Gaps

ö

(first entry) 06-MAY-2004 Oligonucleotide associated to ILSR-X61176 #115.

interleukin; IL-4 receptor; IL-5 receptor; lung disease; airway inflammation; allergy; asthma; impeded respiration; cystic fibrosis; acute respiratory distress syndrome; pulmonary hypertension; lung inflammation; bronchitis; oligonucleotide;

Homo sapiens.

WO2004011613-A2.

05-FEB-2004.

25-JUL-2003; 2003WO-US023509.

29-JUL-2002; 2002US-0399076P.

(EPIG-) EPIGENESIS PHARM INC.

Aguilar D, Miller Nyce JW, Tang L, Sandrasagra A, Shahabuddin S, Lu H, Cong H;

ö

WPI; 2004-203534/19.

e.g., Novel single or multiple target oligonucleotide anti-sense to e.g. initiation codons and introns of respiratory disease-relevant genes CCR1, RANTES, MCP4, useful for prophylaxis or treating respiratory disease e.g., asthma. 

Claim 2; SEQ ID NO 2279; 85pp; English.

initiation codon, coding region with 2-10 nucleotides of 5-end and 3'c interleukin (IL)-4 receptor, IL-5 receptor or salts of the
coligonucleotide and optionally surfactant operatively linked to the
coligonucleotide. The method is useful for preventing or treating a
respiratory or lung disease, which involves administering to the airways
cc of a subject an effective amount of an inhibitor. The oligonucleotide is
useful for production of a medicament for the prevention and/or treatment
c da respiratory or lung disease. The respiratory or lung disease is
chosen from airway inflammation, allergy(ies), asthma, impeded
respiration, cystic fibrosis (CP), chronic obstructive pulmonary diseases
(COPD), allergic rhinitis (AR), acute respiratory distress syndrome
(ARDS), pulmonary hypertension, lung inflammation, bronchitis, airway
cobstruction. The present sequence represents an oligonucleotide of the present invention relates to an oligonucleotide anti-sense to The

10001863-3.sl.rng

Local Similarity

```
The present invention relates to an oligonucleotide anti-sense to e.g., initiation codon, coding region with 2-10 nucleotides of 5'-end and 3'-end of nucleic acid target comprising gene(s) chosen from e.g. interleukin (IL)-4 receptor, IL-5 receptor or salts of the cideral condition of the method is useful for preventing or treating a respiratory or lung disease, which involves administering to the airways of subject an effective amount of an inhibitor. The oligonucleotide is useful for production and/or treatment of a respiratory or lung disease. The respiratory or lung disease. The respiratory or lung disease. The respiratory or lung disease is chosen from alrway inflammation, allergy(ses), asthma, impeded consen from alrway inflammation, allergy(ses), asthma, impeded consen from alrway inflammation, allergy(ses), asthma, impeded consenting chinitis (AR), acute respiratory distress syndrome (ARDS), pulmonary hypertension, lung inflammation, bronchitis, airway obstruction. The present sequence represents an oligonucleotide of the
                                                                                                                                                                                                            ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Novel single or multiple target oligonucleotide anti-sense to e.g. initiation codons and introns of respiratory disease-relevant genes e.g., CCR1, RANTES, MCP4, useful for prophylaxis or treating respiratory
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  airway inflammation; allergy; asthma; impeded respiration;
cystic fibrosis; acute respiratory distress syndrome;
pulmonary hypertension; lung inflammation; bronchitis; oligonucleotide;
                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                            ;
0
                                                                                                                                        Length 20;
                                                                                                                                                                                                        3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              interleukin; IL-4 receptor; IL-5 receptor; lung disease;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Miller S;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 20 BP; 5 A; 6 C; 5 G; 4 T; 0 U; 0 Other;
                                                                    Sequence 20 BP; 9 A; 6 C; 3 G; 2 T; 0 U; 0 Other;
                                                                                                                                    Score 15.2; DB 1;
Pred. No. 2.3e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Aguilar D,
                                                                                                                                                                                                    0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Oligonucleotide associated to MCP4 #25.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Claim 2; SEQ ID NO 763; 85pp; English.
                                                                                                                                                                                                                                                                        2873 AAAATACAGAGTCTTCCAGG 2892
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Nyce JW, Tang L, Sandrasagra A,
Shahabuddin S, Lu H, Cong H;
                                                                                                                                                                                                                                                                                                                                         1 AAAACACAGAATCCTCCAGG 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     H.
                                                                                                                                    0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               25-JUL-2003; 2003WO-US023509.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   29-JUL-2002; 2002US-0399076P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (EPIG-) EPIGENESIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (first entry)
                                                                                                                                    Query Match 0.4 Best Local Similarity 85.0 Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ADJ59907 standard; DNA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    disease e.g., asthma.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2004-203534/19.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WO2004011613-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         06-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              05-FEB-2004.
invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ADJ59907;
                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 30

ADJ5 9907/
TO ADJ5 9907/
TO ADJ5 9907/
XX XX XX XX ADJ5 9907/
XX XX XX ADJ5 9907/
XX ADJ5
   SXS
                                                                                                                                                                                                                                                                        ò
                                                                                                                                                                                                                                                                                                                                     셤
```

DB 1; Length 20;

0.4%; Score 15.2;

Query Match

```
The present invention relates to an oligonucleotide anti-sense to e.g., initiation codon, coding region with 2-10 nucleotides of 5'-end and 3'-end of nucleic acid target comprising gene(s) chosen from e.g. interleukin (IL)-4 receptor, IL-5 receptor or salts of the cligonucleotide and optionally surfactant operatively linked to the oligonucleotide. The method is useful for preventing or treating a respiratory or lung disease, which involves administering to the airways of a subject an effective amount of an inhibitor. The oligonucleotide is useful for production of a medicament for the prevention and/or treatment of a respiratory or lung disease. The respiratory or lung disease. The respiratory or lung disease. The respiratory or lung disease is chosen from airway inflammation, allergy(iss), asthma, impeded respiration, cystic fibrosis (CF), chronic obstructive pulmonary diseases (CPDD), allergic rhinitis (AR), acute respiratory distress syndrome (CDDD), allergic rhinitis (AR), acute respiratory bronchitis, airway obstruction. The present sequence represents an oligonucleotide of the
                  ô
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              e.g.,
                                                                                                                                                                                                                                                                                                                                            interleukin; IL-4 receptor; IL-5 receptor; lung disease;
airway inflammation; allergy; asthma; impeded respiration;
cystic fibrosis; acute respiratory distress syndrome;
pulmonary hypertension; lung inflammation; bronchitis; oligonucleotide;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                    Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Novel single or multiple target oligonucleotide anti-sense to e.g. initiation codons and introns of respiratory disease-relevant genes CCR1, RANTES, MCP4, useful for prophylaxis or treating respiratory disease e.g., asthma.
                    ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 2.3e+02; Matches 17; Conservative 0; Mismatches 3; Indels
                Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ŝ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Miller
85.0%; Pred. No. 2.3e+02; ive 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 5 A; 3 C; 8 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Aguilar D,
                                                                                                                                                                                                                                                                                                        Oligonucleotide associated to ICAM #195.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Claim 2; SEQ ID NO 1277; 85pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         3750 TTTAGGGAGACACAGATGGC 3769
                                                      3184 TCTCCTTACAGAGGTTAAAG 3203
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              L, Sandrasagra A,
Lu H, Cong H;
                                                                                            20 rcrccrrccacacaccrcaac
                                                                                                                                                                                          ADJ60421 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              25-JUL-2003; 2003WO-US023509.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    29-JUL-2002; 2002US-0399076P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        (EPIG-) EPIGENESIS PHARM INC.
                                                                                                                                                                                                                                                                    (first entry)
                  17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2004-203534/19.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WO2004011613-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Nyce JW, Tang
Shahabuddin S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens.
                                                                                                                                                                                                                                                                  06-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          05-FEB-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        .nvention.
                                                                                                                                                                                                                              ADJ60421;
                                                                                                                                                    RESULT 301
                  Matches
                                                                                                                                                                        ADJ60421
                                                        ò
                                                                                          g
                                                                                                                                                                                            ò
```

RESULT 303

1 TTGAGGGGGACACAGATGTC 20

셤

```
New compounds, particularly antisense oligonucleotides targeted to a
nucleic acid encoding resistin, useful for treating a metabolic disorder,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                The invention describes a compound 8-80 nucleobases in length targeted to, and which specifically hybridises with a nucleic acid molecule encoding resistin, and inhibits the expression of resistin. The compound, composition and methods are useful for treating a disease or condition associated with resistin, such as a metabolic disease, e.g. diabetes or obesity, or atherosclerosis. They are also useful in research and diagnostics for modulating the expression of resistin. This sequence
                                                                                                                                                                                                                                                                                         antidiabetic, anorectic, cardiant, antiarteriosclerotic, resistin, inhibitor, resistin; metabolic disease, diabetes, obesity, atherosclerosis, antisense technology, human; antisense oligonucleotide,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  /mod_base= OTHER /mosphorothioate backbone. All cytidines /note= "OTHER= Phosphorothioate backbone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         /mod_base= OTHER
/note= "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /*tag= c
/mod_base= OTHER
/note= "OTHER= 2'-0-Methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Score 15.2; DB 1; Length 20;
Pred. No. 2.38+02;
0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        represents a human resistin antisense oligonucleotide
                                                                                                                                                                                                                                Human resistin antisense oligonucleotide seg id 111.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 20 BP; 4 A; 3 C; 9 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             nucleic acid encoding resistin, useful for tr
e.g. diabetes or obesity, or atherosclerosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Example 15; SEQ ID NO 111; 75pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            31-JUL-2002; 2002US-00210833.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               31-JUL-2002; 2002US-00210833
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1. .5
/*tag= a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          *tag= b
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               .20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             17; Conservative
                                 ADJ38722 standard; DNA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2004-142664/14.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   US2004023383-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  modified base
                                                                                                                                                                 06-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Bhanot S,
                                                                                                 ADJ38722;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Matches
Abujastzz

Abujastz

Abujas
```

```
The invention relates to a method for treating airway hyperresponsiveness or pulmonary inflammation in an individual comprising administering an antisense compound targeted to a nucleic acid molecule encoding a human B7 protein. The invention also relates to a method of inhibiting expression of a nucleic acid molecule encoding B7-1 or B7-2. The antisense compound is an antisense oligonucleotide which has a modified sugar molecy and nucleobase. The human B7 protein is human B7-1 or B7-2 protein or both. The compound is useful for treating airway hyperresponsiveness or pulmonary inflammation, which is associated with asthma, by inhibiting expression of human B7 protein. This sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Treating airway hyperresponsiveness or pulmonary inflammation comprises administering an antisense compound targeted to a nucleic acid molecule encoding a human B7 protein to the individual.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    B7 protein. This sequence the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ö
                                                                                                                                 Airway hyperresponsiveness; pulmonary inflammation; antisense oligonucleotide; human; B7 protein; B7-1; asthma; antiasthmatic; antiinflammatory; 8s.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.4%; Score 15.2; DB 1; Length 20;
35.0%; Pred. No. 2.36+02;
lve 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 20 BP; 3 A; 4 C; 4 G; 9 T; 0 U; 0 Other;
                                                                                                 Human B7-1 DNA antisense oligonucleotide #33.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           represents an antisense oligonucleotide of
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Example 12; SEQ ID NO 128; 182pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Karras JG;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1317 TIGAGITICAAAGGITGCIG 1336
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1 TTTAGTTTCACAGCTTGCTG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   BP
                                                                                                                                                                                                                                                                                                         23-MAY-2003; 2003US-00444206.
                                                                                                                                                                                                                                                                                                                                       31-DEC-1996; 96US-00777266.
04-JUN-1999; 99US-00326186.
25-MAY-2000; 2000WO-US014471.
09-MAY-2001; 2001US-00851871.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             85.0%;
ADJ54308 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADJ54433 standard; DNA; 20
                                                               (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      06-MAY-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Bennett CF, Vickers TA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                          (BENN/) BENNETT C F. (VICK/) VICKERS T A.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2004-132608/13.
                                                                                                                                                                                                                                                                                                                                                                                                                                            (VICK/) VICKERS T A (KARR/) KARRAS J G.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Local Similarity
                                                                                                                                                                                                                                       US2004023917-A1.
                                                                                                                                                                                                      Homo sapiens.
                                                               06-MAY-2004
                                                                                                                                                                                                                                                                       05-FEB-2004
                               ADJ54308
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADJ54433;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 30-
ADJ54433/
               ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   · B
```

ö

ö

Gaps ö

547 TACAGAAGCTGGTGGCTGTG 566

ઠ d

1 racadgaaccccicrcrc 20

```
The invention relates to a method for treating airway hyperresponsiveness or pulmonary inflammation in an individual comprising administering an antisense compound targeted to a nucleic acid molecule encoding a human B7 protein. The invention also relates to a method of inhibiting a human expression of a nucleic acid molecule encoding B7-1 or B7-2. The antisense compound is an antisense oligomucleotide which has a moified sugar moiety and nucleobase. The human B7 protein is human B7-1 or B7-2 protein or both. The compound is useful for treating airway propensiveness or pulmonary inflammation, which is associated with asthma, by inhibiting expression of human B7 protein. This sequence represents an antisense oligonucleotide of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Antisense therapy; mouse; complement component C3; autoimmune disorder; multiple sclerosis; infection; atherosclerosis; neuroprotective; antiarteriosclerotic; antimicrobial; antiinflammatory; cytostatic; phosphorothioate; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Treating airway hyperresponsiveness or pulmonary inflammation comprises administering an antisense compound targeted to a nucleic acid molecule encoding a human B7 protein to the individual.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Mouse complement component C3 DNA, antisense oligonucleotide #51.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%; Score 15.2; DB 1; Length 20; 35.0%; Pred. No. 2.3e+02; ive 0; Mismatches 3; Indels
                                                              antisense oligonucleotide; human; B7 protein; B7-1; asthma; antiasthmatic; antiinflammatory; ss.
                                               Airway hyperresponsiveness; pulmonary inflammation;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 7 A; 4 C; 8 G; 1 T; 0 U; 0 Other;
              Human B7-1 DNA antisense oligonucleotide #88
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Example 19; SEQ ID NO 253; 182pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                    Bennett CF, Vickers TA, Karras JG;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      3767 GGCTGGGATCCCTCCCTGT 3786
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     20 GGCTGGCATCCCTCTCCTTT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ADK12305 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                           31-DEC-1996; 96US-0077266.
04-UTN-1999; 99US-00326186.
25-MAY.-2000; 2000WO-US014411.
09-MAY-2001; 2001US-00851871.
                                                                                                                                                                                                                          23-MAY-2003; 2003US-00444206
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 82.08;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     17; Conservative
                                                                                                                                                                                                                                                                                                                                                  (BENN/) BENNETT C F. (VICK/) VICKERS T A. (KARR/) KARRAS J G.
                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2004-132608/13.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Local Similarity
                                                                                                                                                        US2004023917-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   20-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Mus musculus
                                                                                                                         Homo sapiens.
                                                                                                                                                                                           05-FEB-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ADK12305;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT 305
ADK12305
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Best Loca
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ð
```

ó

Gaps

. 0

```
The present invention relates to antisense compounds targeted to a nucleic acids encoding human and mouse complement component C3. The antisense compound comprises an antisense oligonucleotide that specifically hybridises with the nucleic acid and inhibits the expression of complement component C3 in cells. The antisense oligonucleotide is a chimeric oligonucleotide. The antisense oligonucleotide is a chimeric oligonucleotide. The antisense oligonucleotide comprises at chast one modified internucleoside linkage, preferably a phosphorothhoate linkage. It also comprises at least one modified augar moiety, preferably a 2'-0-methoxyethyl (2'-MOB) sugar moiety. The antisense oligonucleotide further comprises at least one modified nucleobase, preferably a 5-cmethylytosine. The antisense oligonucleotide is an entisense such as autocimmune disorders e.g. multiple creament of diseases such as autocimmune disorders e.g. multiple selezosis, infections, and atherosclerosis. The present sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ô
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Antilipaemic; Cardiovascular; Analgesic; Antianginal; Antisense therapy; Human; Endothelial Lipase; dyslipidaemia; high density lipoprotein; \operatorname{HDL}; cardiovascular disorder; \operatorname{metabolic} syndrome X_i ss.
                                                                /mcd_base= OTHER
/mode= "This oligonucleotide has a phosphorothioate
/note= "This oligonucleotide has a the 5'
backbone and 2'-methyoxyethyl (2'-MOE) wings at the 5'
and 3' ends, which are 5 nucleotides in length at each
end. All cytidine residues are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                        New antisense compound targeted to a nucleic acid molecule encoding complement component C3, useful for treating multiple sclerosis, an infection or atherosclerosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Human endothelial lipase antisense oligonucleotide, SEQ ID 2658.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           .
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match

0.4%; Score 15.2; DB 1; Length 20;
Best Local Similarity 85.0%; Pred. No. 2.3e+02;
Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Seguence 20 BP; 5 A; 6 C; 4 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Claim 3; SEQ ID NO 163; 74pp; English.
             Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1399 TTACCATGAGTTCAAACTTC 1418
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ВР
                                                                                                                                                                                                                                                          18-AUG-2003; 2003US-00642802.
                                                                                                                                                                                                                                                                                                23-OCT-2001; 2001US-00001076.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADJ24260 standard; DNA; 20
                               1. .20
/*tag= a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (first entry)
                                                                                                                                                                                                                                                                                                                                                                                              Graham MJ, Watt AT;
                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2004-225730/21
                                                                                                                                                                                                                                                                                                                                   (GRAH/) GRAHAM M J. (WATT/) WATT A T.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 present invention.
                                                                                                                                                                                    US2004043956-A1
                 Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Homo sapiens,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20-MAY-2004
                                                                                                                                                                                                                         04-MAR-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ADJ24260;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 306
ADJ24260/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       셤
```

```
WO2004009541-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Key
modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               20-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                           17;
                                                                       29-JAN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADJ24576;
                                                                                                                                                                                                                                                                                                                                                                                       Query Match
                                                                                                                                                       Bhat BG;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   RESULT 308
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ADJ24576/c
                                                                                                                                                                                                                                                                                                                                                                                                           Matches
 ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                  셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                            New antisense oligonucleotide for modulating endothelial lipase expression, for diagnosing, preventing or treating e.g. dyslipidemia, low high density lipoprotein or cardiovascular disorders.
                                               /mod_base= OTHER
/note= "This oligonucleotide has a phosphorothicate
backbone and 2-'methyoxyethyl (2'-MOE) wings at the 5'
and 3' ends, which are 4 nucleotides in length. Also all
cytidine residues are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Antilipaemic, Cardiovascular, Analgesic, Antianginal, Antisense therapy, Human, Endothelial Lipase; dyslipidaemia, high density lipoprotein, HDL, cardiovascular disorder, metabolic syndrome X; ss.
                                                                                                                                                                                                                                                                                                                       The present invention relates to antisense oligonucleotides (ADJ21603-ADJ25510) targeted to human Endothelial Lipase (EL) coding sequence (ADJ25517), where the antisense oligonucleotide specifically hybridises with and inhibits the expression of EL. The antisense oligonucleotides are useful for modulating the expression of endothelial lipase in cells or tissues to treat diseases associated with EL expression, such as dyslipidaemia, low high density lipoprotein (HDL), cardiovascular disorder or metabolic syndrome X. In addition, the oligonucleotides are used for diagnostics, prophylaxis, or as research reagents or kits.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Human endothelial lipase antisense oligonucleotide, SEQ ID 47.
                                                                                                                                                                                                                                                                                                                                                                                                                                                0.4%; Score 15.2; DB 1; Length 20; 15.0%; Pred. No. 2.3e+02; ve 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                Sequence 20 BP; 10 A; 3 C; 2 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                   Claim 3; SEQ ID NO 2658; 1007pp; English
                    Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1897 AGAATGACTTTGCTTGTACT 1916
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        20 AGAATGATTTTTTTTGTACT 1
                                                                                                                                                       18-JUL-2003; 2003WO-US022410
                                                                                                                                                                            19-JUL-2002; 2002US-0397106P
                                                                                                                                                                                                                                                                                                                                                                                                                                                             85.0%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ADJ21649 standard; DNA; 20
                                          æ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                            Local Similarity 85.0
                                                                                                                                                                                              (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                         WPI; 2004-132912/13.
                                                                                                              WO2004009541-A2
                    Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            20-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Bapiens
                                                                                                                                   29-JAN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Synthetic.
 Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ADJ21649;
                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                                                                                                                                    Bhat BG;
                                                                                                                                                                                                                                                                                                                                                                                                                                                               Best Loca
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Ношо
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 30
ADJ21649
ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            g
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 New antisense oligonucleotide for modulating endothelial lipase expression, for diagnosing, preventing or treating e.g. dyslipidemia, low high density lipoprotein or cardiovascular disorders.
/note= "This oligonucleotide has a phosphorothioate backbone and 2-"methyoxyethyl (2-'MCB' wings at the 5' and 3' ends, which are 4 nucleotides in length. Also all cytidine residues are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Antilipaemic; Cardiovascular; Analgesic; Antianginal; Antisense therapy; Human; Endothelial Lipase; dyslipidaemia; high density lipoprotein; HDL; cardiovascular disorder; metabolic syndrome X; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /mod_base= OTHER
/note= "This oligonucleotide has a phosphorothioate
backbone and 2-'methyoxyethyl (2'-MOE) wings at the 5'
and 3' ends, which are 4 nucleotides in length. Also all
cytidine residues are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             The present invention relates to antisense oligonucleotides (ADJ21603-ADJ25510) targeted to human Endothelial Lipase (EL) coding sequence (ADJ25517), where the antisense oligonucleotide specifically hybridises with and inhibits the expression of EL. The antisense oligonucleotides are useful for modulating the expression of endothelial lipase in cells or tissues to treat diseases associated with EL expression, such as dyslipidaemia, low high density lipoprotein (HDL), cardiovascular disorder or metabolic syndrome X. In addition, the oligonucleotides are used for diagnostics, prophylaxis, or as research reagents or kits.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human endothelial lipase antisense oligonucleotide, SEQ ID 2974
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      3; Indele
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 20 BP; 4 A; 5 C; 5 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Claim 3; SEQ ID NO 47; 1007pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Location/Qualifiers
1. .20
/*tag= a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                3130 ACTGCAGTCGTCAAGGGGCT 3149
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ACTGCAGTCTTCTAAGGGCT 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ВP
                                                                                                                                                                                                                                                                                                                                             18-JUL-2003; 2003WO-US022410.
                                                                                                                                                                                                                                                                                                                                                                                                                   19-JUL-2002; 2002US-0397106P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ADJ24576 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           WPI; 2004-132912/13.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Local Similarity
                                                                                                                                                                                            WO2004009541-A2.
```

10001863-3.sl.rng

```
New antisense compound targeted to a nucleic acid molecule encoding Nav1.3, useful for useful for treating a disease or condition associated with Nav1.3, e.g. pain, seizure disorder such as childhood seizure
                  18-JUL-2003; 2003WO-US022410
                               19-JUL-2002; 2002US-0397106P
                                                                                                                                                                                                                                                                                                                                                                                         14-AUG-2003; 2003WO-US025465.
                                                                                                                                                                                                                                                                                                                                                                                                     14-AUG-2002; 2002US-0403416P.
                                           (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                  (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2004-203785/19.
                                                                    WPI; 2004-132912/13.
                                                                                                                                                                                                        Local Similarity
                                                                                                                                                                                                                                                                                                                                                               WO2004016754-A2
                                                                                                                                                                                                                                                                                                 20-MAY-2004
      29-JAN-2004
                                                                                                                                                                                                                                                                                                                                                                            26-FEB-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                               Roberds SL;
                                                                                                                                                                                                                                                                                                                                                   Synthetic.
                                                                                                                                                                                                                                                                                    ADK79428;
                                                                                                                                                                                                                            1898
                                                                                                                                                                                                  Query Match
                                                        Bhat BG;
                                                                                                                                                                                                              Matches
                                                                                                                                                                                                                                                           RESULT 309
                                                                                                                                                                                                                                                                ò
                                                                                                                                                                                                                                      셤
```

; 0 New antisense oligonucleotide for modulating endothelial lipase expression, for diagnosing, preventing or treating e.g. dyslipidemia, low high density lipoprotein or cardiovascular disorders. The present invention relates to antisense oligonucleotides (ADJ21603-ADJ25510) targeted to human Endothelial Lipase (EL) coding sequence (ADJ25517), where the antisense oligonucleotide specifically hybridises with and inhibits the expression of EL. The antisense oligonucleotides are useful for modulating the expression of endothelial lipase in cells or tissues to treat diseases associated with EL expression, such as dyslipidaemia, low high density lipoprotein (HDL), cardiovascular disorder or metabolic syndrome X. In addition, the oligonucleotides are used for diagnostics, prophylaxis, or as research reagents or kits. Gaps ; 0 Length 20; 3; Indels Sequence 20 BP; 11 A; 3 C; 2 G; 4 T; 0 U; 0 Other; Score 15.2; DB 1; Pred. No. 2.3e+02; 0; Mismatches Claim 3; SEQ ID NO 2974; 1007pp; English GAATGACTTTGCTTGTACTT 1917 20 GAATGATTTTTTTTTTTT 1 0.4%; 428/c ADK79428 standard; DNA; 20 17; Conservative

BP.

ADK78831 standard; DNA; 20

ADK7883:

ADK78831;

20-MAY-2004 (first entry)

Navl.3; Analgesic; Nootropic; Neuroprotective; post-herpetic neuralgia; diabetic neuropathy; arthritic pain; migraine headache; infantile epilepsy; ataxia; ss. Chimeric phosphorothioate oligonucleotide to target Nav1.3 #6762. (first entry)

The present invention relates to an antisense compound targeted to a nucleic acid molecule encoding Navi.3, where the antisense compound specifically hybridizes with and inhibits the expression of Navi.3. The compound and composition are useful for treating a disease or condition associated with Navi.3. e.g. pain including but not limited to neuropathic pain, post-herpetic neuralgia, chronic pain, lower back pain, diabetic neuropathy, trigeminal neuropathy, arthritic pain, acute pain, pain from burns, migraine headache, cluster headache, mild-to-moderate headache; seizure disorder such as childhood seizure disorder, including but not limited to neonatal or infantile epilepsy, or ataxia. The present cepresents a chimeric phosphorothicate oligonucleotide with 2.MOB wings and a deoxy gap. Used during the antisense inhibition of human Navi.3 expression, the oligonucleotides are designed to target Gaps ö Match 0.4%; Score 15.2; DB 1; Length 20; Local Similarity 85.0%; Pred. No. 2.3e+02; les 17; Conservative 0; Mismatches 3; Indels Sequence 20 BP; 3 A; 5 C; 1 G; 11 T; 0 U; 0 Other; Claim 4; SEQ ID NO 6762; 417pp; English. 967 ACTIGGAAAGTITGACAAA 986 20 ACAAGGAAAAGTTTGAGAAA 1 disorder, or ataxia. Query Match RESULT 310 Matches Š 셤

ö

New antisense compound targeted to a nucleic acid molecule encoding Navl.3, useful for useful for treating a disease or condition associated with Navl.3, e.g. pain, seizure disorder such as childhood seizure Navl.3; Analgesic; Nootropic; Neuroprotective; post-herpetic neuralgia; diabetic neuropathy; arthritic pain; migraine headache; Chimeric phosphorothioate oligonucleotide to target Nav1.3 #6165. infantile epilepsy; ataxia; ss. 14-AUG-2002; 2002US-0403416P. 14-AUG-2003; 2003WO-US025465. (PHAA ) PHARMACIA CORP WPI; 2004-203785/19. WO2004016754-A2. 26-FEB-2004. Roberds SL; Synthetic.

The present invention relates to an antisense compound targeted to a nucleic acid molecule encoding Navl.3, where the antisense compound specifically hybridizes with and inhibits the expression of Navl.3. The compound and composition are useful for treating a disease or condition associated with Navl.3, e.g. pain including but not limited to neuropathic pain, post-herpetic neuralgia, chronic pain, lower back pain, 

Claim 4; SEQ ID NO 6165; 417pp; English.

disorder, or ataxia.

```
pain from burns, migraine headache, cluster headache, mild-to-moderate headache; seizure disorder such as childhood seizure disorder, including but not limited to neonatal or infantile epilepsy; or ataxia. The present sequence represents a chimeric phosphorothioate oligonucleotide with human Navl.3 expression, the oligonucleotide with human Navl.3 expression, the oligonucleotides are designed to target different regions of the human Navl.3 RNA.
arthritic pain, acute pain,
diabetic neuropathy, trigeminal neuropathy,
```

8888888888

Sequence 20 BP; 4 A; 5 C; 3 G; 8 T; 0 U; 0 Other;

```
ö
     Length 20;
                          3; Indels
  Score 15.2; DB 1;
Pred. No. 2.3e+02;
0; Mismatches 3;
                                                  1080 TGTTTGACAAATGTTTCTTC 1099
                                                                        1 rerrreaccaarerarcrec 20
0.4%;
            Local Similarity 85.0 nes 17; Conservative
  Query Match
                          Matches
                                                  ઠે
                                                                       셤
```

ö

Gaps

ADK73906 standard; DNA; 20 RESULT 311 

BP

20-MAY-2004 (first entry) ADK73906;

Chimeric phosphorothioate oligonucleotide to target Nav1.3 #1240.

Navl.3; Analgesic; Nootropic; Neuroprotective; post-herpetic neuralgia; diabetic neuropathy; arthritic pain; migraine headache; infantile epilepsy; ataxia; ss.

Synthetic.

WO2004016754-A2

26-FEB-2004.

14-AUG-2003; 2003WO-US025465.

14-AUG-2002; 2002US-0403416P.

(PHAA ) PHARMACIA CORP

Roberds SL;

New antisense compound targeted to a nucleic acid molecule encoding abav1.3, useful for useful for treating a disease or condition associated with Nav1.3, e.g. pain, selzure disorder such as childhood seizure disorder, or ataxia. WPI; 2004-203785/19

Claim 4; SEQ ID NO 1240; 417pp; English.

Included a compound and inhibits the antisense compound specifically hybridizes with and inhibits the expression of Navl.3. The compound and composition are useful for treating a disease or condition associated with Navl.3, e.g. pain including but not limited to neuropathic pain, post-herpetic neuralgia, chronic pain, lower back pain, diabetic neuropathy, trigeminal neuropathy, archritic pain, acute pain, pain from burns, migraine headache, cluster headache, mild-to-moderate headache; seizure disorder such as childhood seizure disorder such but not limited to neonatal or infantile epilepsy; or ataxia. The present but not limited to neonatal or infantile epilepsy; or ataxia. The presequence represents a chimeric phosphorothioate oligonucleotide with 2'MOE wings and a decxy gap. Used during the antisense inhibition of human Navl.3 expression, the oligonucleotides are designed to target different regions of the human Navl.3 RNA. present invention relates to an antisense compound targeted to a

ô

Gaps

ö

0.4%; Score 15.2; DB 1; Length 20; 15.0%; Pred. No. 2.3e+02; ve 0; Mismatches 3; Indels

82.08;

17; Conservative

Matches

ò

- E

Local Similarity

Query Match

940 TTCTGGGAGAATTTAGAAAT 959

1 Tranggagaarrangaaar 20

Sequence 20 BP; 8 A; 0 C; 5 G; 7 T; 0 U; 0 Other;

Sequence 20 BP; 5 A; 2 C; 4 G; 9 T; 0 U; 0 Other;

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             The present invention relates to an antisense compound targeted to a mucleic acid molecule encoding Nav1.3, where the antisense compound specifically hybridizes with and inhibits the expression of Nav1.3. The compound and composition are useful for treating a disease or condition neuropathic pain, post-herpetic neuralgia, chronic pain, lower back pain, encodetic neuropathy, trigeminal neuropathy, arthritic pain, acute pain, pain from burns, migraine headache, cluster headache, mild-to-moderate pain from burns migraine headache, cluster headache, mild-to-moderate but not limited to neonatal or infantile epilepsy, or ataxia. The present sequence represents a chimeric phosphorothioate oligonucleotide with
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          New antisense compound targeted to a nucleic acid molecule encoding about 3, useful for useful for treating a disease or condition associated with Navl.3, e.g. pain, seizure disorder such as childhood seizure disorder, or ataxia.
                                                                                                                                                                                                                                                                                                                                  Navl.3; Analgesic; Nootropic; Neuroprotective; post-herpetic neuralgia; diabetic neuropathy; arthritic pain; migraine headache; infantile epilepsy; ataxia; ss.
                                    Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            2'MOB wings and a deoxy gap. Used during the antisense inhibition of human Navl.3 expression, the oligonucleotides are designed to target different regions of the human Navl.3 RNA.
                                                                                                                                                                                                                                                                                                   Chimeric phosphorothioate oligonucleotide to target Nav1.3 #4894.
                                    ö
   Length 20;
                                  Indele
0.4%; Score 15.2; DB 1;
85.0%; Pred. No. 2.3e+02;
ive 0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Claim 4; SEQ ID NO 4894; 417pp; English.
                                                                     256 AGGTGGTTCCTAATATTACT 275
                                                                                                  1 AGGIGGITACTACTATTATT 20
                                                                                                                                                                                            뗦.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                14-AUG-2003; 2003WO-US025465.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  14-AUG-2002; 2002US-0403416P.
                                                                                                                                                                                          ADK77560 standard; DNA; 20
                                                                                                                                                                                                                                                                (first entry)
                  Local Similarity 85.0 nes 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WPI; 2004-203785/19.
                                                                                                                                                                                                                                                                                                                                                                                                                                            WO2004016754-A2.
                                                                                                                                                                                                                                                                20-MAY-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               26-FEB-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Roberds SL;
                                                                                                                                                                                                                             ADK77560;
   Query Match
                    Best Loca
Matches
                                                                                                                                                         RESULT 312
                                                                       ò
                                                                                                  셤
```

```
Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 315
ð
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              엄
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                              The present invention relates to an antisense compound targeted to a nucleic acid molecule encoding Nav1.3, where the antisense compound specifically hybridizes with and inhibits the expression of Nav1.3. The compound and composition are useful for treating a disease or condition neuropathle pain, post-herpetic neuralgia, chronic pain, lower back pain, diabetic neuropathy, trigeminal neuropathy, arthritic pain, acute pain, pain from burns, migraine headache, cluster headache, mild-to-moderate but not limited to monatal or infantile epilepsy, or ataxia. The present sequence represents a chimeric phosphorothioate oligonuclectide with
                                                                                                                                                                                                                                                                                                                                   New antisense compound targeted to a nucleic acid molecule encoding Navl.3, useful for useful for treating a disease or condition associated with Navl.3, e.g. pain, seizure disorder such as childhood seizure disorder, or ataxia.
                                                                                                          Navl.3; Analgesic; Nootropic; Neuroprotective; post-herpetic neuralgia; diabetic neuropathy; arthritic pain; migraine headache; infantile epilepsy; ataxia; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        2'MOE wings and a deoxy gap. Used during the antisense inhibition of human Navl.3 expression, the oligonucleotides are designed to target different regions of the human Navl.3 RNA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Chimeric phosphorothioate oligonucleotide to target Nav1.3 #4162.
                                                                                     Chimeric phosphorothioate oligonucleotide to target Navl.3 #8938
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 10 A; 4 C; 5 G; 1 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15.2; DB 1; 35.0%; Pred. No. 2.3e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                          Claim 4; SEQ ID NO 8938; 417pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      485 AATATTGACAGGAAACCCCA 504
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AATAGAGACAGGAAAGCCCA 20
                                                                                                                                                                                                                        14-AUG-2003; 2003WO-US025465.
                                                                                                                                                                                                                                               14-AUG-2002; 2002US-0403416P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    82.08;
                  ADK81604 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADK76828 standard; DNA; 20
                                                                (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20-MAY-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                17; Conservative
                                                                                                                                                                                                                                                                  (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                               WPI; 2004-203785/19.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Best Local Similarity
                                                                                                                                                                            WO2004016754-A2
                                                               20-MAY-2004
                                                                                                                                                                                                  26-FEB-2004.
                                                                                                                                                                                                                                                                                          Roberds SL;
                                                                                                                                                       Synthetic.
                                          ADK81604;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 314
ADK76828
ID ADK768
AC ADK768
XX
DT 20-MAY
XX
XX
XX
XX
XX
XX
XX
XX
XX
```

```
The present invention relates to an antisense compound targeted to a mucleic acid molecule encoding Nav1.3, where the antisense compound specifically hybridizes with and inhibits the expression of Nav1.3. The compound and composition are useful for treating a disease or condition neuropaths the pain including but not limited to neuropaths pain, post-herpetic neuralgia, chronic pain, lower back pain, diabetic neuropathy, trigeminal neuropathy, arthritic pain, acute pain, pain from burns, migraine headache, cluster headache, mild-to-moderate but not limited to neonatal or infantile epilepsy, or ataxia. The present sequence represents a chimeric phosphorothioate oligonuclevide with
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     New antisense compound targeted to a nucleic acid molecule encoding Nav1.3, useful for useful for treating a disease or condition associated with Nav1.3, e.g. pain, seizure disorder such as childhood seizure disorder, or ataxia.
Navl.3; Analgesic; Nootropic; Neuroprotective; post-herpetic neuralgia; diabetic neuropathy; arthritic pain; migraine headache; infantile epilepsy; ataxia; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Navl.3; Analgesic; Nootropic; Neuroprotective; post-herpetic neuralgia; diabetic neuropathy; arthritic pain; migraine headache; infantile epilepsy; ataxia; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           2'MOE wings and a deoxy gap. Used during the antisense inhibition of human Navl.3 expression, the oligonucleotides are designed to target different regions of the human Navl.3 RNA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Chimeric phosphorothioate oligonucleotide to target Nav1.3 #1129.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
iive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 20 BP; 5 A; 3 C; 4 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Claim 4; SEQ ID NO 4162; 417pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1078 ATTGTTTGACAAATGTTTCT 1097
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADK73795 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                            14-AUG-2003; 2003WO-US025465.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Best Local Similarity op.v
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2004-203785/19.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Local Similarity
                                                                                                                                                                                                                                        WO2004016754-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WO2004016754-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         14-AUG-2002;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 20-MAY-2004
                                                                                                                                                                                                                                                                                                                    26-FEB-2004,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              26-FEB-2004,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Roberds SL;
```

ö

```
Chimeric phosphorothioate oligonucleotide to target Nav1.3 #1745.
                                                                                                                                                                                                                                      Sequence 20 BP; 4 A; 2 C; 5 G; 9 T; 0 U; 0 Other;
                                                                                                          Claim 4; SEQ ID NO 1129; 417pp; English
                                                                                                                                                                                                                                                                                1074 TITAATIGITIGACAAAIGI 1093
                                                                                                                                                                                                                                                                                             1 TTTGAGTGTTTGACCAATGT 20
                                                                                                                                                                                                                                                                                                                                                                                                            infantile epilepsy; ataxia; ss
14-AUG-2003; 2003WO-US025465
              14-AUG-2002; 2002US-0403416P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                    14-AUG-2003; 2003WO-US025465.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   14-AUG-2002; 2002US-0403416P.
                                                                                                                                                                                                                                                                                                                                    ADK74411 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                20-MAY-2004 (first entry)
                                                                                                                                                                                                                                                  Query Match
Best Local Similarity 85.0 Matches 17; Conservative
                            (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (PHAA ) PHARMACIA CORP
                                                        WPI; 2004-203785/19.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2004-203785/19
                                                                                                                                                                                                                                                                                                                                                                                                                                        WO2004016754-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                      26-FEB-2004.
                                           Roberds SL;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Roberds SL;
                                                                                                                                                                                                                                                                                                                                                                                                                          Synthetic.
                                                                                                                                                                                                                                                                                                                                                  ADK74411;
                                                                                                                                                                                                                                                                                                                    RESULT 316
ADK74411
ID ADK7441
ò
                                                                                                                                                                                                                                                                                             요
```

Score 15.2; DB 1; Pred. No. 2.3e+02; 0; Mismatches

85.0%;

BP

```
nucleic acid molecule encoding Navi. 3, where the antisense compound specifically hybridizes with and inhibits the expression of Navi. 3. The compound and composition are useful for treating a disease or condition associated with Navi. 3, e.g. pain including but not limited to neuropathic pain, post-herpetic neuralgia, chronic pain, lower back pain, diabetic neuropathy, trigeminal neuropathy, arthritic pain, acute pain, pain from burns, migraine headache, cluster headache, mild-to-moderate headache; seizure disorder such as childhood seizure disorder, including but not limited to neonatal or infantile epilepsy; or ataxia. The present sequence represents a chimeric phosphorothioate olisonucletide with
                    Navl.3, useful for useful for treating a disease or condition associated with Navl.3, e.g. pain, seizure disorder such as childhood seizure disorder, or ataxia.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Lipford G;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           2'MOE wings and a deoxy gap. Used during the antisense inhibition of human Nav1.3 expression, the oligonucleotides are designed to target different regions of the human Nav1.3 RNA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                immunostimulatory nucleic acid; asthma; allergy; cancer; infectious disease; autoimmune disease; airway remodeling; chronic obstructive pulmonary disease; asthma; II-6; interleukin-6; TNFalpha; tumour necrosis factor alpha; IFNalpha; interferon-alpha; IFNalpha; interferon-gamma; IP-10; interferon inducible protein; viral infection; bacteria infection; parasitic infection; ss.
New antisense compound targeted to a nucleic acid molecule encoding
                                                                                                                                                                                                                                                                     The present invention relates to an antisense compound targeted to
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
ative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Jurk M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 20 BP; 5 A; 2 C; 5 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Uhlmann E,
                                                                                                                                                                                    Claim 4; SEQ ID NO 1745; 417pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Samulowitz U, Vollmer J,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Immunostimulatory nucleic acid #93
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     254 GGAGGTGGTTCCTAATATTA 273
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1 Gráddrichachacharha 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (COLE-) COLEY PHARM GROUP INC. (COLE-) COLEY PHARM GMBH.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ADK19046 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          19-AUG-2002; 2002US-0404479P.
19-AUG-2002; 2002US-0404820P.
27-NOV-2002; 2002US-0429701P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      19-AUG-2003; 2003WO-US025935.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   14-FEB-2003; 2003US-0447377P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       20-MAY-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WPI; 2004-257200/24.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WO2004016805-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              26-FEB-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Krieg AM,
Rankin R;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADK19046;
a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      HANNER BY SERVICE STANKER SERVICE STANKER SERVICE STANKER SERVICE STANKER SERVICE STANKER SERVICE SERV
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           The present invention relates to an antisense compound targeted to a nucleic acid molecule encoding Nav1.3, where the antisense compound specifically hybridizes with and inhibits the expression of Nav1.3. The compound and composition are useful for treating a disease or condition associated with Nav1.3, e.g. pain including but not limited to neuropathic pain, post-herpetic neuralgia, chronic pain, lower back pain, diabetic neuropathy, trigeninal neuropathy, arthritic pain, acute pain, pain from burns, migraine headache, cluster headache, mild-to-moderate headache; seizure disorder such as childhood seizure disorder, including but not limited to neonatal or infantile epilepsy; or ataxia. The present sequence represents a chimeric phosphorothioate oligonuclectide with
                                                                                                                                                                                                                                                                                                                                                                             New antisense compound targeted to a nucleic acid molecule encoding abav1.3, useful for useful for treating a disease or condition associated with Nav1.3, e.g. pain, seizure disorder such as childhood seizure disorder, or ataxia.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Navl.3; Analgesic; Nootropic; Neuroprotective; post-herpetic neuralgia; diabetic neuropathy; arthritic pain; migraine headache;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             sequence represents a chimeric phosphorothioate oligonucleotide with 2'MOS wings and a deoxy gap. Used during the antisense inhibition of human NaV1.3 expression, the oligonucleotides are designed to target different regions of the human NaV1.3 RNA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              3; Indels
```

New immunostimulatory nucleic acid molecule having pyrimidine-purine dinucleotide and a chimeric backbone, useful in treating and preventing

ö

Page 156

10001863-3.sl.rng

```
The invention relates to an immunostimulatory nucleic acid molecule comprising an internal pyrimidine-purine (YZ) dinucleotide and chimeric backbone, where one internal YZ dinucleotide has a phosphodiester(-like) internal variational internal YZ dinucleotide ach additional internal YZ dinucleotide linkages are stabilised. The cligonucleotide linkages are stabilised. The cligonucleotide is useful in stimulating or modulating an immune response. The medicament shifts the immune response to a Thi biased response. The oligonucleotide is also useful in the manufacture of a medicament for treating as thin allergy, cancer, intectious disease, autoimmune disease, airway remodeling or chronic obstructive pulmonary disease or in treating a subject who is a smoker or who is free of symptoms of asthma. The oligonucleotide is useful in corrective pulmonary disease or in treating a subject who is a smoker or who is free of symptoms of asthma. The oligonucleotide is useful in correction and in treating and preventing infections caused by viruses, also useful in treating and preventing infections caused by viruses, immunication and preventing infections caused by viruses, immunication; and paresters and preventing infections caused by viruses, immunication; imminications caused by viruses, immunication; imminications caused by viruses, immunication; imminications caused by viruses.
asthma, allergy, cancer, infectious disease, autoimmune disease or airway remodeling.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Human; 88; interleukin-4 receptor; IL-4; interleukin-5 receptor; IL-5; CCR1; CCR3; Eccaxin-1; RANTES; MCP4; CD23; ICAM; UCAM; tryptase a; tryptase b; PD84 B; PD84 C; PD84 D; respiratory disease; lung disease; hyper-responsiveness; adenosine; adenosine A receptor; asthma; lung allergy; inflammation; inflammatory disease; airway inflammation; allergy; impeded respiration; cystic fibrosis; CF; chronic obstructive pulmonary disease; COPD; allergic rhinitis; acute respiratory distress syndrome; pulmonary hypertension; lung inflammation; bronchitis; airway obstruction; bronchoconstriction.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%; Score 15.2; DB 1; Length 20; 15.0%; Pred. No. 2.3e+02; ve 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 20 BP; 0 A; 2 C; 2 G; 16 T; 0 U; 0 Other;
                                                              Claim 4; SEQ ID NO 93; 276pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        3326 TITICATITITIACGICT 3345
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1 rrrcgrrrrrrrcgrrrr 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              immunostimulatory nucleic acid.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ADO45397 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             25-JUL-2003; 2003US-00627930
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        23-APR-2002; 2002WO-US013135.
23-APR-2002; 2002WO-US013143.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      82.08;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human oligonucleotide #763.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     15-JUL-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
Best Local Similarity 85.0°
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          US2004049022-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   11-MAR-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AD045397;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 318
ADO45397/c
  ઠે
```

NYCE J W. SANDRASAGRA A. TANG L. AGUILAR D. MILLER S. SHAHABUDDIN S.

(NYCE/) R (SAND/) R (TANG/) R (AGUI/) R (MILL/) R

```
The invention relates to oligonucleotides anti-sense to an initiation codon, coding region, 5' or 3' intron-exon junction, intron or region with 2-10 nucleotides of the 5'-end or 3'-end of a nucleic acid target chosen from a gene encoding interleukin (IL)-4 receptor, interleukin (IL) crossen from a gene encoding interleukin (IL)-4 receptor, interleukin (IL) crossen from a gene encoding interleukin (IL)-4 receptor, interleukin (IL) crossen from a gene encoding interleukin (IL)-4 receptor, interleukin (IL) crossen from a gene encoding darget(s) or expressed product(s), for the prevention and/or treatment of a respiratory or lung disease. The coligonucleotides are useful for reducing or inhibiting expression of a coligonucleotides are useful for reducing or inhibiting expression of a crospiratory or lung disease. The coligonucleotides are useful for reducing or inhibiting expression of a crospiratory or lung disease is associated with hyper-responsiveness to and/or increased levels of, adenosine and/or levels of adenosine A creeptor(s), and/or asthma and/or lung allergie associated with crossing colificammation or an inflammation, allergy, asthma, impeded respiration, crossing (CP), chronic obstructive pulmonary disease (COPD), allergic rhinitis, acute respiratory distress syndrome, pulmonary crossing crossing crospiratory distress syndrome, pulmonary crossing 
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ö
                                                                                                                                                                                                                                                                                                           Novel single or multiple target oligonucleotide anti-sense to e.g. initiation codon, intron of respiratory disease-relevant gene e.g. CCR1, RANTES, MCP4, useful for prophylaxis or treating respiratory disease e.g.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human; ss; interleukin-4 receptor; IL-4; interleukin-5 receptor; IL-5; CCR1; CCR3; Ectaxin-1; RANTES; MCP4; CD23; ICAM; VCAM; tryptase a; tryptase b; PD24 A; PDE4 C; PDE4 C; PDE4 C; PDE4 C; PDE5 C; 
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 2.3e+02; Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                             Aguilar D, Miller S;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Sequence 20 BP; 5 A; 6 C; 5 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Claim 2; SEQ ID NO 763; 174pp; English.
                                                                                                                     Tang L,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    3184 TCTCCTTACAGAGGTTAAAG 3203
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20 rcrccrrccacacacaca
                                                                                Sandrasagra A, Tany
Tan H, Cong H;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ADO45910 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human oligonucleotide #1276.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             15-JUL-2004 (first entry)
                                                                                                                                                     Shahabuddin S, Lu H,
                                                                                                                                                                                                                                      WPI; 2004-293804/27.
(CONG/) LU H.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ADO45910;
                                                                                                                     Nyce JW,
                                                                                                                                                                                                                                                                                                                                                                                                                                     asthma.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RESULT 319
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               AD045910
à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             셤
```

```
25-JUL-2003; 2003US-00627930
                      23-APR-2002; 2002WO-US013135.
23-APR-2002; 2002WO-US013143.
                                 NYCE J W.
SANDRASAGRA A.
                                                MILLER S.
SHAHABUDDIN S.
                                                                              WPI; 2004-293804/27
                                         TANG L.
AGUILAR D.
US2004049022-A1
                                                            CONG/) CONG H.
       11-MAR-2004.
                                                                                                                                                                                                        invention
                                                                    Nyce JW,
                                                    (SHAH/)
(LUHH/)
                                 NYCE/)
                                             AGUI/)
                                         TANG/)
                                                 MILL/)
                                     (SAND/)
                                                                                                  asthma
```

Sequence 20 BP; 5 A; 3 C; 8 G; 4 T; 0 U; 0 Other;

3750 TTTAGGGAGACACAGATGGC 3769

ADO46813 standard; DNA; 20 BP. ADO46813; RESULT 320
ADO46813
ID ADO4681
XX
AC ADO4681
XX
DT 15-JUL-

15-JUL-2004 (first entry)

Novel single or multiple target oligonucleotide anti-sense to e.g. initiation codon, intron of respiratory disease-relevant gene e.g. CCR1, RANTES, MCP4, useful for prophylaxis or treating respiratory disease e.g. A, Tang L, Aguilar D, Miller S; Cong H; Claim 2; SEQ ID NO 1277; 174pp; English. Sandrasagra A, Shahabuddin S, Lu H,

The invention relates to oligonucleotides anti-sense to an initiation codon, coding region, 5' or 3' intron-exon junction, intron or region with 2-10 mucleotides of the 5'-end or 3'-end of a nucleic acid target chosen from a gene encoding interleukin (IL)-4 receptor, interleukin (IL)-5 receptor, CCR1, CCR3, Ectaxin-1, RANTES, MCP4, CD23, ICAM, VCAM, Tryptase a, tryptase b, PDE4 A, PDE4 B, PDE4 C or PDE4 D. The invention also relates to a method of screening a candidate compound that binds to one or more nucleic acid target(8) or expressed product(8), for the prevention and/or treatment of a respiratory or lung disease. The coligonucleotides are useful for reducing or inhibiting expression of a gene or mRNA encoding interleukin-5 receptor, CCR1, CCR3, Eotaxin-1, RANTES, MCP4, CD23, ICAM, VCAM, tryptase a, pDE4 A, PDE4 B, PDE4 C, or PDE4 D. The oligonucleotides are useful for preventing or treating a respiratory or lung disease. The respiratory or lung disease is associated with hyper-responsiveness to and/or increased levels of, adenosine and/or levels of adenosine A inflammation or an inflammation, allergy, asthma, impeded respiration, cleaned from airway inflammation, allergy, asthma, impeded respiration, allergic fibrosis (CP), chronic obstructive pulmonary or lung disease is considered with inflammation, allergy, asthma, impeded respiration, hypertension, lung inflammation, bronchitis, airway obstruction or hymerican. This sequence represents an oligonucleotide of the

Gaps ö Length 20; 3; Indels Score 15.2; DB 1; Pred. No. 2.3e+02; 0; Mismatches 3; 0.4%; Query Match
Best Local Similarity 85.0
Matches 17; Conservative

ö

|| |||| ||| ||||||||| || TTGAGGGGGACACAGATGTC 20

유 ò

Query Match

Human oligonucleotide #2179.

Human; 88; interleukin-4 receptor; IL-4; interleukin-5 receptor; IL-5; CCR1; CCR3; Eccaxin-1; RANTES; MCP4; CD21; ICAM; VCAM; tryptase a; tryptase b; PD84 B; PD84 C; PD84 D; respiratory disease; lung disease; hyper-responsiveness; adenosine; adenosine A receptor; asthma; lung allergy; inflammation; inflammatory disease; alway inflammation; allergy; impeded respiration; cystic fibrosis; CF; alrway inflammation; allergy; disease; COPD; allergic rhinitis; acute respiratory diserses syndrome; pulmonary hypertension; lung inflammation; bronchitis; airway obstruction; bronchoconstriction. 

Homo sapiens.

US2004049022-A1.

11-MAR-2004.

25-JUL-2003; 2003US-00627930.

23-APR-2002; 2002WO-US013135. 23-APR-2002; 2002WO-US013143.

NYCE J W. SANDRASAGRA A. TANG L. AGUILAR D. (NYCE/) (TANG/) MILL/

MILLER S. SHAHABUDDIN S.

CONG/) CONG H. (SHAH/) (LUHH/)

Nyce JW, Sandrasagra A, Tang L, Aguilar D, Miller S; Shahabuddin S, Lu H, Cong H;

WPI; 2004-293804/27.

Novel single or multiple target oligonucleotide anti-sense to e.g. initiation codon, intron of respiratory disease-relevant gene e.g. CCR1, RANTES, MCP4, useful for prophylaxis or treating respiratory disease e.g. asthma.

Claim 2; SEQ ID NO 2279; 174pp; English.

The invention relates to oligonuclectides anti-sense to an initiation codon, coding region, 5' or 3' intron-exon junction, intron or region with 2-10 nucleotides of the 5'-end or 3'-end of a nucleic acid target chosen from a gene encoding interleukin (1L)-4 receptor, interleukin (1L) coseptor, cCR1, CCR3, Eotaxin-1, RANTES, MCP4, CD23, ICAM, VCAM, LTYptase a, tryptase b, PDE4 A, PDE4 B, PDE4 C or PDE4 D. The invention also relates to a method of screening a candidate compound that binds to one or more nucleic acid target(s) or expressed product(s), for the prevention and/or treatment of a respiratory or lung disease. The coligonucleotides are useful for reducing or inhibiting expression of a gene or mRNA encoding interleukin-4 receptor, interleukin-5 receptor, CCR1, CCR3, Botaxin-1, RANTES, MCP4, CD23, ICAM, VCAM, tryptase a, CCR1, CCR3, Botaxin-1, RANTES, MCP4, CD23, ICAM, VCAM, tryptase a, creptratory or lung disease is associated with hyper-responsiveness to useful for preventing or treating a respiratory or lung disease. The creptor(s), and/or asthma and/or lung allergies associated with creased levels of adenosine A creceptor(s), and/or asthma and/or lung allergies associated with creased levels of adenosine A creceptor(s), and/or asthma and/or lang allergy, asthma, impeded respiration, cystic. fibrosis (CF), chronic obstructive pulmonary disease (COPD), allergic rhinitis, acute respiratory discress syndrome, pulmonary disponention or hypertension, lung inflammation, bronchiefs, and allergy and allergic rhinitis, acute respiratory discress syndrome, pulmonary disponention or hypertension, lung inflammation, bronchiefs, and decomposite and allergic rhinitis, acute respiratory discress syndrome, pulmonary disponention or hypertension, allergic rhinitis, acute respiratory discress syndrome, pulmonary disponention or hypertension, lung inflammation, bronchiefs, and decomposite and decomposited ronchoconstriction. This sequence represents an oligonucleotide of the

Sequence 20 BP; 9 A; 6 C; 3 G; 2 T; 0 U; 0 Other;

0.4%; Score 15.2; DB 1; Length 20;

ö

```
The invention describes a compound 8-80 nucleobases in length targeted to a nucleic acid molecule encoding Cytokine-inducible kinase. The compound specifically hybridises with the nucleic acid molecule encoding Cytokine-inducible kinase (which comprises a sequence of 2169 bp fully defined in the specification) and inhibits the expression of Cytokine-inducible kinase (which comprises a sequence of 1869 bp fully defined in Cytokine-inducible are: a method of inhibiting the expression of Cytokine-inducible kinase in cells or tissues, comprising contacting the cells or tissues with the new compound so that the expression of Cytokine-inducible kinase is inhibited; a method of screening for a modulator of Cytokine-inducible kinase, comprising contacting a preferred target segment of the nucleic acid encoding Cytokine-inducible kinase with one or more candidate modulators of Cytokine-inducible kinase with one
                                                                                                                                                                                                                                                                                                                     cytokine-inducible kinase inhibitor; antisense technology; cytokine-inducible kinase expression; hyperproliferative disorder; human; antisense oligonucleotide; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                             /*tag= b
/mod_base= OTHER
/note= "OTHER= Phosphorothioate backbone. All cytidines
are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      New antisense oligonucleotides useful for modulating Cytokine-inducible kinase expression, useful for diagnosing, preventing or treating conditions associated with aberrant kinase expression e.g. hyperproliferative disorders.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /mod_base= OTHER
/note= "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              /mod_base= OTHER
/note=_"OTHER= 2'-0-Methoxyethyl (2'-MOE) nucleotides"
                                Gaps
                                ;
0
                                                                                                                                                                                                                                                                               Human cytokine-inducible kinase antisense oligonucleotide #17.
                                                                                                                                                                                                                                                                                                                cytostatic; antisense therapy; cytokine-inducible kinase;
                                Indels
. 2.3e+02;
3;
                              0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Example 15; SEQ ID NO 32; 56pp; English.
                 Pred. No.
                                                                                                                                                                                                                                                                                                                                                                                                                                Location/Qualifiers
                                                               2873 AAAATACAGAGTCTTCCAGG 2892
                                                                                              1 AAAACACAGAATCCTCCAGG 20
                                                                                                                                                                               BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               23-NOV-2002; 2002US-00304116.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                23-NOV-2002; 2002US-00304116.
             82.0%;
                                                                                                                                                                             ADN30046 standard; DNA; 20
                                                                                                                                                                                                                                             12-AUG-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   υ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              /*tag= a
             Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             15. .20
/*tag= c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2004-399685/37.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Ward DT, Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               US2004101857-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              modified base
                                                                                                                                                                                                                                                                                                                                                                                               Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                27-MAY-2004
                                                                                                                                                                                                              ADN30046;
                                                                                                                                                RESULT 321
                                                                                                                                                                ADN30046/
                                                               à
                                                                                           셤
```

```
identifying one or more modulators that modulate the expression of
Cytokine-inducible kinase, a diagnostic method for identifying a disease
state, comprising identifying the presence of Cytokine-inducible kinase
in a sample using at least one of the primers or probe comprising the
nucleotide sequences as mentioned in the specification; a kit or assay
device comprising the above compound, and a method of treating an animal
having a disease or condition associated with Cytokine-inducible kinase,
comprising administering to the animal a therapeutic or prophylactic
amount of the compound so that expression of Cytokine-inducible kinase in
inhibited. The antisense oligonucleotide is useful for inhibiting the
expression of Cytokine-inducible kinase in cells or tissues to prevent or
treat diseases associated with the kinase expression, such as
hyperproliferative discorders. In addition, the compound is used for
diagnostics, prophylaxis, or as research reagents or kits. This sequence
represents a human cytokine-inducible kinase antisense oligonucleotide.
                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           cytostatic, antisense therapy, cytokine-inducible kinase, cytokine-inducible kinase inhibitor; antisense technology; cytokine-inducible kinase expression; hyperproliferative disorder; human;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /mod_base= OTHER /note= "OTHER= Phosphorothioate backbone. All cytidines are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      15. 20
/*tage c
//mod base= OTHER
/note= "OTHER= 2'-0-Methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Human cytokine-inducible kinase antisense oligonucleotide #87.
                                                                                                                                                                                                                                                                                                            Query Match 0.4%; Score 15.2; DB 1; Length 20; Best Local Similarity 85.0%; Pred. No. 2.3e+02; Matches 17; Conservative 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                Sequence 20 BP; 5 A; 9 C; 1 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                    399 GAACTGCAGGTGCTGGATTT 418
                                                                                                                                                                                                                                                                                                                                                                                                                    20 GAACTGAAGGTGGGGGATTT 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 antisense oligonucleotide; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ADN30116 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               23-NOV-2002; 2002US-00304116.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             23-NOV-2002; 2002US-00304116.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           12-AUG-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Ω
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        *tag= a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ...20
*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WPI; 2004-399685/37.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Ward DT, Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            US2004101857-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              27-MAY-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ADN30116;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 322
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADN30116
       8$88888888888888
                                                                                                                                                                                                                                                                                                                                                                                    ઠ
                                                                                                                                                                                                                                                                                                                                                                                                                   셤
```

10001863-3.81.rng

```
The interior describes a compount of the interior action of the compound specifically hybridises with the nucleic acid molecule encoding Cytokine-inducible kinase. The compound inhibits the expression of Cytokine-inducible in the specification) and inhibits the expression of Cytokine-inducible kinase (which comprisis comprising contacting the expression of Cytokine-inducible kinase in cells or tissues, comprising contacting the colls or tissues with the new compound so that the expression of Cytokine-inducible kinase, comprising contacting a preferred target segment of the nucleic acid encoding Cytokine-inducible kinase, and compression of Cytokine-inducible kinase, comprising contacting a preferred target segment of the nucleic acid encoding Cytokine-inducible kinase, and compression of Cytokine-inducible kinase, and compression of cytokine-inducible kinase, and dentifying the presence of Cytokine-inducible kinase is take, comprising identifying the presence of Cytokine-inducible kinase is take, comprising the above compound; and method of treating an animal device comprising the above compound; and a method of treating an animal device comprising the above compound; and a method of treating an animal computation as of the animal a therapeutic or prophylactic amount of the antisense oligonucleotide is useful for inhibiting the expression of Cytokine-inducible kinase is inhibited. The antisense oligonucleotide is useful for inhibiting the expression of cytokine-inducible kinase is contracted and the cytokine-inducible kinase is comprising administering the difference of cytokine-inducible kinase is comprising administering the difference of cytokine-inducible kinase is comprising administering the difference of comprision of cytokine-inducible kinase is comprised to that expression of Cytokine-inducible kinase is comprised to the compound of the antisense of cytokine-inducible kinase is comprised to the compound to the animal and the cytokine-inducible kinase is confired to the cytokine-inducible kinase is conf
                                                                                                                                                                                     The invention describes a compound 8-80 nucleobases in length targeted to
New antisense oligonucleotides useful for modulating Cytokine-inducible kinase expression, useful for diagnosing, preventing or treating conditions associated with aberrant kinase expression e.g.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /note= "OTHER= Phosphorothioate backbone. All cytidines are 5-methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            /mod_base= OTHER /note= "OTHER= 2'-0-Methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      represents a human cytokine-inducible kinase antisense oligonucleotide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      treat diseases associated with the kinase expression, such as hyperproliferative disorders. In addition, the compound is used for diagnostics, prophylaxis, or as research reagents or kits. This sequ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       cytostatic; antisense therapy; oestrogen-responsive finger protein; oestrogen-responsive finger protein associated disorder; hyperproliferative disorder; diagnostic; prophylaxis; human; antisense oligonucleotide; antisense technology; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Oestrogen-responsive finger protein antisense oligo seqid 72
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; ive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 20 BP; 5 A; 1 C; 9 G; 5 T; 0 U; 0 Other;
                                                                                                                              Example 15; SEQ ID NO 102; 56pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     399 GAACTGCAGGTGCTGGATTT 418
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    GAACTGAAGGTGGGGGATTT 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           BP
                                                                             hyperproliferative disorders.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ADP31773 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1. .20
/*tag= b
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /*tag= a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Local Similarity 85.0
Les 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 26-AUG-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADP31773;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       RESULT 323
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    셤
```

```
The invention describes a compound 8-80 nucleobases in length targeted to a nucleic acid molecule encoding oestrogen-responsive finger protein. The compound specifically hybridises with the nucleic acid molecule encoding cestrogen-responsive finger protein (which comprises a sequence of 24295 bp fully defined in the specification) and inhibits the expression of estrogen-responsive finger protein. Also described are: a method of cinhibiting the expression of cestrogen-responsive finger protein, in cells or tissues; a method of screening for a modulator of oestrogen-responsive finger protein; a diagnostic method for identifying a disease state; a kit or assay device comprising the above compound; and a method of treating an animal having a disease or condition associated with estrogen responsive finger protein. The antisense oligonucleotide is useful for inhibiting the expression of oestrogen-responsive finger protein in cells or tissues to prevent or treat diseases associated with aberrant contissues tinger protein expression, such as hyperproliferative disorders. In addition, the compound is used for diagnostics, prophylaxis, or as research reagents or kits. This sequence represents a human oestrogen-responsive finger protein antisense
                                                                                                                                                                                                                                                                                                                                                                    New antisense oligonuclectides for modulating estrogen-responsive finger protein expression, useful for diagnosing, preventing or treating hyperproliferative disorders.
                                      /mod_base= OTHER
/note= "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   cytostatic; antisense therapy; oestrogen-responsive finger protein; oestrogen-responsive finger protein associated disorder; hyperproliferative disorder; diagnostic; prophylaxis; human; antisense oligonuclectide, antisense technology; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Oestrogen-responsive finger protein antisense oligo seqid 147
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; tive 0; Mismatches 3; Indel8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 3 A; 5 C; 4 G; 8 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                   Example 15; SEQ ID NO 73; 65pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         2479 AGAAGGTGGAGAAGACCCTG 2498
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ATAACGTGGAGAAAACCCTG 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADP31848 standard; DNA; 20 BP
                                                                                                                                                                            10-DEC-2002; 2002US-00317277.
                                                                                                                                                                                                                  10-DEC-2002; 2002US-00317277.
                    /*tag= c
/mod_base= (
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          26-AUG-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   17; Conservative
                                                                                                                                                                                                                                                         (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                 WPI; 2004-440347/41.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match
Best Local Similarity
                                                                                               US2004110159-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   oligonucleotide
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Homo sapiens
                                                                                                                                       10-JUN-2004
                                                                                                                                                                                                                                                                                               Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      RESULT 324
ADP31848
    ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            원
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Gaps
```

ö

Gaps

```
The invention relates to a compound, having a sequence comprising 8-80 bp targeted to a nucleic acid encoding BAF53 (a member of the BAF complex (BRG1/brm-associated factor 53kDa which is an actin-related protein), specifically hybridises with the nucleic acid encoding BAF53 comprising 28001-bp sequence (derived from human chromosome 3) and inhibits expression of BAF53, i.e. an antisense oligonucleotide. Also included are inhibiting the expression of BAF53 in cells or tissues, screening for a modulator of BAF53, a diagnostic method for identifying a disease state, a kit or assay device comprising the compound and treating an animal having a disease or condition associated with BAF53. The oligonucleotide compound is useful for preparing a tumour. The present sequence is an antisense oligonucleotide targeting
                                                       Human; ss; antisense; chromatin; BAF complex; BRG1/brm-associated factor; BAF53; BRG1-associated factor 53kDa; cancer; tumour; actin-related protein; hyperproliferative disorder.
                                                                                                                                                                                                                                     /note= "Phosphorothioate backbone and all cytidines are methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     New oligonucleotide compound that inhibits expression of BAF53, useful for preparing a composition for treating hyperproliferative disorder,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Score 15.2; DB 1;
Pred. No. 2.3e+02;
0; Mismatches 3;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 20 BP; 2 A; 7 C; 2 G; 9 T; 0 U; 0 Other;
                  Human BAF53 antisense oligonucleotide ISIS280335.
                                                                                                                                                                                                                                                                                                                                                                                                                    /*tag= a
/mod_base= OTHER
/note= "2'-methoxyethyl_residue"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Example 15; SEQ ID NO 29; 72pp; English
                                                                                                                                                                                              Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             3438 ACTCTGTCACTTTGTCACTC 3457
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Acrerereagrirererere 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DEC-2002; 2002US-00316243.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                09-DEC-2002; 2002US-00316243.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ADP49620/c
ID ADP49620 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                     1. .20
/*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WPI; 2004-440336/41
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Jain R;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  US2004110147-A1.
                                                                                                                                                                                                                   modified base
                                                                                                                                                                                                                                                                                                                                 modified base
                                                                                                                                                                                                                                                                                                                                                                                                                        modified base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   numan BAF53
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             10-JUN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Dobie KW,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         RESULT 326
                                                                                                                                                       Ношо
ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     The invention describes a compound 8-80 nucleobases in length targeted to a nucleic acid molecule encoding oestrogen-responsive finger protein. The compound specifically hybridises with the nucleic acid molecule encoding oestrogen-responsive finger protein (which comprises a sequence of 24295 bp fully defined in the specification) and inhibits the expression of estrogen-responsive finger protein. Also described are: a method of inhibiting the expression of oestrogen-responsive finger protein; a method of screening for a modulator of oestrogen-responsive finger protein; a diagnostic method for identifying a disease state; a kit or assay device comprising the above compound; and a method of treating an animal having a disease or condition associated with estrogen responsive finger protein. The antisense oligonucleotide is useful for inhibiting the expression of oestrogen-responsive finger protein. The antisense oligonucleotide is useful for inhibiting the expression of oestrogen-responsive finger protein in cells or tissues to prevent or treat diseases associated with aberrant
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         oestrogen_responsive finger protein expression, such as typerproliferative disorders. In addition, the compound is used for diagnostics, prophylaxis, or as research respents or kits. This sequence represents a human oestrogen-responsive finger protein antisense
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              estrogen-responsive finger
                                                         /mod_base= OTHER
/note= "OTHER= Phosphorothioate backbone. All cytidines
are 5-methylcytidines"
                                                                                                                                                                       /mod_base= OTHER
/note= "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides"
15. .20
                                                                                                                                                                                                                                                                                   /note= "OTHER= 2'-O-Methoxyethyl (2'-MOE) nucleotides
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       New antisense oligonucleotides for modulating estrogen-responsive protein expression, useful for diagnosing, preventing or treating hyperproliferative disorders.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; ive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Sequence 20 BP; 8 A; 4 C; 5 G; 3 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Example 15; SEQ ID NO 148; 65pp; English.
Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          2479 AGAAGGTGGAGAAGACCCTG 2498
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /mod_base= OTHER
                                                                                                                                                                                                                                                                                                                                                                                                                        10-DEC-2002; 2002US-00317277
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  LO-DEC-2002; 2002US-00317277
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20
                                                                                                                                                       *tag= a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    26-AUG-2004 (first entry)
                                                                                                                                                                                                                                              Ų
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Best Local Similarity 85.0
Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADP49542 standard; DNA;
                                                                                                                                                                                                                                              /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 2004-440347/41
                                                                                                                                                                                                                                                                                                                                 US2004110159-A1
Key
modified_base
                                                                                                                              modified_base
                                                                                                                                                                                                                     modified_base
                                                                                                                                                                                                                                                                                                                                                                             10-JUN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ADP49542;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Dobie KW
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
```

ö

Gaps

ö

Length 20; Indels

ò

```
Human; d8; antisense; chromatin; BAF complex; BRG1/brm-associated factor; BAF53; BRG1-associated factor 53kDa; cancer; tumour; actin-related protein; hyperproliferative disorder; chromosome 3.
                                                                                                                                                                                                                                          New oligonucleotide compound that inhibits expression of BAF53, useful for preparing a composition for treating hyperproliferative disorder, e.g. cancer.
                                             Human BAF53 antisense oligonucleotide target region #10.
                                                                                                                                                                                                                                                                                          Example 15; SEQ ID NO 107; 72pp; English.
                                                                                                                                                          09-DEC-2002; 2002US-00316243
                                                                                                                                                                             09-DEC-2002; 2002US-00316243
                         26-AUG-2004 (first entry)
                                                                                                                                                                                              (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                    WPI; 2004-440336/41.
                                                                                                                                                                                                                  Jain R;
                                                                                                                     US2004110147-A1.
                                                                                                    Homo sapiens.
                                                                                                                                        10-JUN-2004.
                                                                                                                                                                                                                 Dobie KW,
        ADP49620;
```

ö The invention relates to a compound, having a sequence comprising 8-80 bp targeted to a nucleic acid encoding BAF53 (a member of the BAF complex (BRG1/brm-associated factor). BRG1-associated factor 53kpa which is an actin-related protein), specifically hybridises with the nucleic acid encoding BAF53 comprising 28001-bp sequence (derived from human chromosome 3) and inhibits expression of BAF53, i.e. an antisense oligonuclectide. Also included are inhibiting the expression of BAF53 in cells or tissues, screening for a modulator of BAF5, a diagnostic method for identifying a disease state, a kit or assay device comprising the compound and treating an animal having a disease or condition associated with BAF53. The oligonuclectide compound is useful for preparing a composition for treating hyperproliferative disorder, e.g. cancer or a tumnour. The BAF53 gene is located on chromosome 3. The present sequence is a BAF53 genenic DNA target sequence for an antisense oligonucleotides. Gaps ö 0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; tive 0; Mismatches 3; Indels Sequence 20 BP; 9 A; 2 C; 7 G; 2 T; 0 U; 0 Other; Best Local Similarity Query Match

3438 ACTCTGTCACTTTGTCACTC 3457 Matches 17; Conservative

20 ACTCTGTCAGTTTCTCTCTC 1

ઠ 셤

ADP49161 standard; DNA; 20 BP. 

ADP49161;

Human nicotinic receptor alpha 7 antisense nested PCR primer.

09-SEP-2004 (first entry)

ss; proinflammatory cytokine; appendicitis; cholinergic agonist; alpha-7 nicotinic receptor; antiinflammatory; gastrointestinal; antilucer; hepatotropic; virucide; antiasthmatic; antiallergic; antibacterial; immunosuppessive; vasotropic; vulnerary; antipyretic;

antimularial, antianginal; cardiant; antiateriosclerotic; thrombolytic; antianualizatal, antianginal; cardiant; antiateriosclerotic; thrombolytic; ophthalmological; cytostatic; osteopathic; antianthritic; ophthalmological; nephrotropic; uropathic; nootropic; antidiabetic; dermatological; nephrotropic; uropathic; nootropic; antidiabetic; local inflammatory disease; urogenital system; respiratory system; infection; dermatological disease; system; respiratory system; cardiovascular system disorder; central nervous system; peripheral nervous system; bone; joint; muscle; connective tissue; autoimmune disorder; inflammatory disorder; cancer; tumour; proliferative disorder; primer; nested PCR; nicotinic receptor; alpha 1 subunit. 

Homo sapiens.

WO2004052365-A2.

24-JUN-2004.

05-DEC-2003; 2003WO-US038708.

06-DEC-2002; 2002US-0431650P

(NSHO-) NORTH SHORE-LONG ISLAND JEWISH RES.

Tracey KJ, Wang H;

WPI; 2004-468700/44.

Treatment of a condition e.g. allergy mediated by release of proinflammatory cytokine involves treating a patient with a cholinergic agonist selective for an alpha-7 nicotinic receptor to decrease the released amount of the cytokine.

Example 1; Page 40; 75pp; English

The inversion relates to a novel method for treatment of a patient cutfering from a condition mediated by release of proinflammatory cytokine e.g. appendicitis involving treating a patient with a cholinesgic agonist (al) selective for an alpha-7 nicotinic receptor to decrease the amount of the proinflammatory cytokine which is released from a macrophage. A cholinergic agonist has antiinflammatory.

CC decrease the amount of the proinflammatory cytokine which is released from a macrophage. A cholinergic agonist has antiinflammatory.

CC decrease the amount of the proinflammatory cytokine which is released antiallaried, antimalerial, immunosuppressive, vasotropic, vulnerary, antipyretic, immunomodulator, gynaecological, respiratory, CNS, anti-HIV, confider, antimalarial, antianginal, cardiant, antiarteriosclerctic, thrombolytic, antimalarial, antianginal, cardiant, antiarteriosclerctic, antiarthritic, ophthalmological, cytostatic, osteopathic, antigout, antiarthritic, ophthalmological, cytostatic, osteopathic, antigout, antiarthritic, ophthalmological, cytostatic, osteopathic, antigout, antigoriatic activity. A compound of the invention is useful for the treatment of diseases involving the gastrointestinal tract and associated tissues, systemic or local inflammatory diseases and conditions, diseases involving the respiratory system and associated tissues, diseases involving the central or peripheral conditions of the skin, diseases involving the central or peripheral conservative tissues, autoimmune and inflammatory disorders as expense or represents and connective tissues, autoimmune and inflammatory disorders as expense. invention relates to a novel method for treatment of a patient subunit of a nicotinic receptor.

Sequence 20 BP; 6 A; 3 C; 7 G; 4 T; 0 U; 0 Other;

Gaps ö Query Match

0.4%; Score 15.2; DB 1; Length 20;
Best Local Similarity 85.0%; Pred. No. 2.3e+02;
Matches 17; Conservative 0; Mismatches 3; Indels

ö

2476 TGCAGAGGTGGAGAGACC 2495

10001863-3.81.rng

**Page 162** 

```
This invention relates to the identification of a novel susceptibility locus AST-1 for asthma and other IgE mediated diseases mapped to the humann chromosome Toll4-pi5. Specifically, it refers to two overlapping genes namely GPRA (G-protein coupled receptor for asthma susceptibility) and AAA1 (asthma associated alternatively spliced gene 1). The present invention describes identifying single nucleotide polymorphisms, as well as insertion or deletion polymorphisms, occurring at different positions in the AST-1 locus, and furthermore providing vectors, host cells, primers and probes in order to determine the status of an individual. Accordingly, it provides a kit to diagnose or assess prediagosition to asthma, chronic obstructive pulmonary disease or cancer and other IgE mediated diseases including rhinitis and dermatitis, such that derived pharmaceutical compositions exhibit cytostatic and antiasthmatic activities. Furthermore, it provides a transgenic animal comprising the asthma locus-1 (AST-1) DNA. This oligonucleotide sequence is a 5' splice junction of the human AAA1 gene, given in Table 11 of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   New GPRA polypeptides, useful in preparing a composition for diagnosing, treating or preventing asthma, other IgE-mediated disease, chronic obstructive pulmonary disease or cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    G-protein coupled receptor for asthma susceptibility; AAA1; asthma associated alternatively spliced gene 1; chronic obstructive pulmonary disease; cancer; rhinitis; dermatitis; cytostatic; antiasthmatic; transgenic; asthma locus-1.
                                                                                                                                                                                                                                                                                                                                                                                                           5' donor site at the exon 9 splice junction of human AAA1 DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ss; AST-1; asthma; IgE mediated disease; human; GPRA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 20 BP; 4 A; 5 C; 5 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.4%; Score 15.2; DB 1;
85.0%; Pred. No. 2.3e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Polvi A,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Laitinen LA,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Example 7; Page 83; 265pp; English.
1 TGCAGATGATGGTGAAGACC 20
                                                                                                                                                                                   BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         20-DEC-2002; 2002US-0435846P.
03-JAN-2003; 2003US-0437895P.
26-MAR-2003; 2003US-0458767P.
09-UUL-2003; 2003US-0486000P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       19-DEC-2003; 2003WO-FI000973
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Salmikangas P;
                                                                                                                                                                                   ADP84370 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Kere J,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WPI; 2004-500286/47.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (GENE-) GENEOS OY.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WO2004056866-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Laitinen T,           
Pulkkinen  V,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Homo sapiens.
                                                                                                                                                                                                                                                                                                                                    23-SEP-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             08-JUL-2004.
                                                                                                                                                                                                                                                          ADP84370;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Query Match
                                                                                                     RESULT 3
ADP 84370
ADP 84370
ADP 84370
ADP 84370
ADP 84370
BD 85'-39-
BD 86'-30-
BD 86'-
음
```

Vendelin J;

Maekelae S,

```
The invention relates to an antisense compound targeted to a nucleic acid molecule encoding acyl-coenzyme A synthetase 1 (ACSI). The antisense compound specifically hybridises with and inhibits the expression of ACSI. The antisense oligonuclectides or compounds are useful for inhibiting the expression of acyl-coenzyme A synthetase 1 (ACSI), and for treating diseases or conditions associated with aberrant expression of ACSI, e.g. diabetes, obesity, metabolic syndrome X, cardiovascular disorder or cancer. The antisense compounds are also useful as research reagents and kits, or in diagnostic, therapeutic and prophylactic applications, e.g. to prevent or delay infection, inflammation or tunour formation. The present sequence represents an acyl-coenzyme A synthetase
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              New antisense compounds targeted to nucleic acid molecules encoding acyl-
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 coenzyme A synthetase 1 (ACS1), useful for treating diseases or conditions associated with aberrant expression of ACS1, e.g. diabetes,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                  Acyl-coenzyme A synthetase 1, ACS1, antisense oligonucleotide #2734.
                                                                                                                                acyl-coenzyme A synthetase 1; ACS1; diabetes; obesity; metabolic syndrome X; cardiovascular disorder; cancer; infection; inflammation; tumour; antisense; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ..
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      PCR; ss; 20 alpha-hydration steroid dehydrogenase; 20-HSD; progesterone metabolism; animal model; miscarriage; livestock;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Score 15.2; DB 1; Length 20;
Pred. No. 2.3e+02;
0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      PCR primer E2F used to amplify murine 20-HSD DNA Seg 3.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 20 BP; 5 A; 4 C; 6 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Claim 3; SEQ ID NO 2734; 940pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1, ACS1, antisense oligonucleotide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                415
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1 CAAGATCTTCAGGTGCTGGA
ADK22657 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                396 CCAGAACTGCAGGTGCTGGA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ADS12549 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                        14-AUG-2003; 2003WO-US025389.
                                                                                                                                                                                                                                                                                                                                        14-AUG-2002; 2002US-0403591P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%;
                                                                (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Conservative
                                                                                                                                                                                                                                                                                                                                                                            (PHAA ) PHARMACIA CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2004-203782/19.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Local Similarity
tes 17; Conserv
                                                                                                                                                                                                                                        WO2004016749-A2.
                                                                  18-NOV-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        16-DEC-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   obesity or
                                                                                                                                                                                                      Synthetic.
                                 ADK22657;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ADS12549;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Query Match
                                                                                                                                                                                                                                                                                                                                                                                                              Ross SA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    RESULT 330
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     ADS12549/c
                ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          В
```

ö

· 0

Gaps

. 0

3; Indels

0; Mismatches

Best Local Similarity 85.0 Matches 17; Conservative

2375 GTCCCAGCACTTCATCCAGA 2394

8

20 Grccaagcacrrcargaga 1

Length 20;

This invention relates to a novel gene that encodes a protein exhibiting 20 alpha-hydration steroid dehydrogenase (20-HSD) activity, an enzyme involved in progesterone metabolism. Specifically, it refers to a mouse animal model that lacks the gene encoding this enzyme, and hence provides a model that can be used for the study of miscarriage resulting from the progesterone dysbolism of a parent or offspring. The present invention describes progesterone dysbolism as a cause of miscarriage in meal resource animals in the livestock farming industry, such that it provides a route to investigate a treatment method and/ or an early check up method with respect to reproductive disturbances occurring in these animals. This oligonucleotide sequence is a PCR primer given in an exemplification of the invention.

New gene encoding a protein exhibiting 20(alpha)-hydration steroid dehydrogenation enzyme activity, useful for metabolizing progesterone to inactive 20(alpha)-dihydroprogesterone.

Disclosure; SEQ ID NO 12; 43pp; Japanese.

ö

Gaps

; 0

0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02; iive 0; Mismatches 3; Indele

ଚ g

Local Similarity 85.0 les 17; Conservative

Query Match Matches BP

ADT00671 standard; DNA; 20

ADT00671;

Sequence 20 BP; 5 A; 4 C; 2 G; 9 T; 0 U; 0 Other;

```
This invention relates to a novel gene that encodes a protein exhibiting 20 alpha-hydration steroid dehydrogenase (20-HSD) activity, an enzyme involved in progesterone metabolism. Specifically, it refers to a mouse animal model that lacks the gene encoding this enzyme, and hence provides a model that can be used for the study of miscarriage resulting from the progesterone dysbolism of a parent or offspring. The present invention describes progesterone dysbolism as a cause of miscarriage in meal resource animals in the livestock farming industry, such that it provides a route to investigate a treatment method and/ or an early check up method with respect to reproductive disturbances occurring in these animals. This oligonucleotide sequence is a PCR primer given in an
                                                                                                                                                                                                                            New gene encoding a protein exhibiting 20(alpha)-hydration steroid dehydrogenation enzyme activity, useful for metabolizing progesterone to inactive 20(alpha)-dihydroprogesterone.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Sequence 20 BP; 9 A; 2 C; 4 G; 5 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                    Disclosure; SEQ ID NO 3; 43pp; Japanese
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1882 TCTTAAATCTTACTCAGAAT 1901
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           exemplification of the invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           20 TCTTCAATCTTGCTTAGAAT 1
 reproductive dysbolism; primer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              H.
                                                                                                            28-FEB-2003; 2003JP-00054612
                                                                                                                                          28-FEB-2003; 2003JP-00054612
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ADS12558 standard; DNA; 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   16-DEC-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              17; Conservative
                                                                                                                                                                                                 WPI; 2004-665472/65.
                                                                                                                                                                     UYTY ) UNIV TOKYO
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Local Similarity
                                                       JP2004261075-A.
                                                                                   24-SEP-2004
                          Rattus sp.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ADS12558;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 RESULT 331
ADS12558
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Matches
셤
```

```
tyrosine kinase; cancer; anti-cancer agent; signalling molecule; tumourigenesis; somatic alteration; colorectal cancer; NTRK3; FES; GUCY2F; MCCK; MLK4; kinase domain; cytostatic; tyrosine kinase inhibitor;
                                                             Novel mutant protein tyrosine kinase-related oligonucleotide SeqID659
                                                                                                                                                                                                            guanylate cyclase stimulator; ss.
      16-DEC-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 2004-718702/70.
                                                                                                                                                                                                                                                                                                                                WO2004082458-A2.
                                                                                                                                                                                                                                                                         Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                       30-SEP-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Bardelli A,
                                                             ö
                                                             Gaps
                                                             ;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            PCR; ss; 20 alpha-hydration steroid dehydrogenase; 20-HSD; progesterone metabolism; animal model; miscarriage; livestock; reproductive dysbolism; primer.
0.4%; Score 15.2; DB 1; Length 20; 85.0%; Pred. No. 2.3e+02;
                                                          3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     PCR primer E2R used to amplify murine 20-HSD DNA Seg 12.
                                                             0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             JP2004261075-A.
```

```
This invention relates to a novel activated mutant protein tyrosine kinases and associated methods for diagnosing cancer and screening for anti-cancer agents. Protein kinases are signalling molecules involved in tumourigenesis. Mutational analysis of the human tyrosine kinase gene
                                                                                                                                                                                                                                                                                                                                                         Activated mutant protein tyrosine kinases (e.g. NTRK3, FES and MCCK) and associated methods for diagnosing cancer and screening for anti-cancer
                                                                                                                                                                                                                                                                                                           Vogelstein
                                                                                                                                                                                                                                                                                                          Parsons W, Velculescu V, Kinzler KW,
                                                                                                                                                                                                                                                                                                                                                                                                         Disclosure; SEQ ID NO 659; 363pp; English.
                                                                                                                                                                                                                      18-FEB-2004; 2004WO-US004452.
                                                                                                                                                                                                                                             21-FEB-2003; 2003US-0448537P.
29-MAY-2003; 2003US-0473895P.
                                                                                                                                                                                                                                                                                  (UYJO ) UNIV JOHNS HOPKINS
```

28-FEB-2003; 2003JP-00054612 28-FEB-2003; 2003JP-00054612.

24-SEP-2004.

Mus musculus

WPI; 2004-665472/65.

(UYTY ) UNIV TOKYO

Sequence 20 BP; 4 A; 3 C; 8 G; 5 T; 0 U; 0 Other;

```
family identified somatic alteration sin 1 in 5 colorectal cancers, with the majority of mutations occurring in the NTRK3, FES, GUCY2F and MCCK/MLK4 genes. Most were identified in the kinase domain. The invention may be useful for the production of compounds with a cytostatic activity acting as protein tyrosine kinase inhibitors or guanylate cyclase stimulators. The invention may be useful for developing methods for detecting mutations involved in cancer or screening for anti-cancer agents. The present sequence is that of a human-derived oligonucleotide which is related to the invention.
         8888888888888
```

Sequence 20 BP; 1 A; 6 C; 3 G; 10 T; 0 U; 0 Other;

```
Gaps
                            .
0
Length 20;
                           3; Indels
Score 15.2; DB 1;
Pred. No. 2.3e+02;
                         0; Mismatches
                                                   3081 TGTTCACTTTTTCCTTTTTG 3100
                                                                            1 rérecacririricecerrie 20
 0.4%;
85.0%;
                         17; Conservative
             Best Local Similarity
 Query Match
                                                                           셤
```

ö

ВР ADT01064 standard; DNA; 20 Novel mutant protein tyrosine kinase-related oligonucleotide SegID1052

(first entry)

16-DEC-2004

ADT01064;

tyrosine kinase; cancer; anti-cancer agent; signalling molecule; tumourigenesis; somatic alteration; colorectal cancer; NTRK3; FES; GUCY2F; MCCK; MLK4; kinase domain; cytostatic; tyrosine kinase inhibitor; guanylate cyclase stimulator; ss.

Homo sapiens

WO2004082458-A2.

30-SEP-2004.

18-FEB-2004; 2004WO-US004452

21-FEB-2003; 2003US-0448537P, 29-MAY-2003; 2003US-0473895P,

(UYJO ) UNIV JOHNS HOPKINS.

Bardelli A, Parsons W, Velculescu V, Kinzler KW, Vogelstein

WPI; 2004-718702/70

Activated mutant protein tyrosine kinases (e.g. NTRK3, FES and MCCK) as associated methods for diagnosing cancer and screening for anti-cancer

Disclosure; SEQ ID NO 1052; 363pp; English.

This invention relates to a novel activated mutant protein tyrosine kinases and associated methods for diagnosing cancer and screening for anti-cancer agents. Protein kinases are signalling molecules involved in tumourigenesis. Mutational analysis of the human tyrosine kinase gene family identified somatic alteration sin 1 in 5 colorectal cancers, with the majority of mutations occurring in the NTRK3, FES, GUCY2F and MCK/MLK4 genes. Most were identified in the kinase domain. The invention may be useful for the production of compounds with a cytostatic activity acting as protein tyrosine kinase inhibitors or guanylate cyclase stimulators. The invention may be useful for developing methods for detecting mutations involved in cancer or screening for anti-cancer agents. The present sequence is that of a human-derived oligonucleotide which is related to the invention. ADTO1064/ XXX ADTO XXX ADTO XXX ADTO XXX ADTO XXX HOMO XXX HOMO XXX HOMO XXX HOMO XXX HOMO XXX ADTO XXX A

```
This is the nucleotide sequence of the PCR primer used for amplification in the method of invention, involving the treatment of type II diabetes with hepatic nuclear factor 4 (HNF-4). The agonists of normal HNF bicactivity can be used to treat diabetes, e.g. to ameliorate disease symptoms involving insufficient expression of an HNF gene and/or canadiorate disease symptoms involving expression of a mutant HNF gene or overexpression of a normal HNF gene. It is also useful to ameliorate disease symptoms involving expression of a mutant HNF gene or overexpression of a normal HNF gene. It is also useful to ameliorate disease symptoms involving a mutant (non-functional) HNF protein e.g. by administering a therapeutically effective amount of an anti-HNF antibody. Protein that binds to the mutant HNF-4 protein is useful in screening a sessays, e.g. to identify antagonists/agonists of an interaction between the Protein and a binding protein to develop drugs for disease treatment. CD Diabetes mellitus is a common metabolic disorder, and most cases are type II (non-insulin dependent diabetes mellitus (NIDDM)). A major genetic component is implicated, but few genes have been identified Mature onset the mutant HNDF have been identified and the form of the few genes have been identified and the few forms have been identified and the few forms have been incharated.
                                              ô
                                                                                                                                                                                                                                                                                                                                                                                                        PCR; primer, amplification; type II diabetes; agonist; antagonist; hepatic nuclear factor 4; HNF-4; HNF gene; anti-HNF antibody; insulin; diabetes mellitus; mature onset diabetes of the young; MODY; MODYI gene;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Treating type II diabetes involving hepatic nuclear factor 4 - useful, e.g. to treat insufficient HNF expression or bioactivity, overexpression of HNF or expression of mutant HNF gene in diabetic patients.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 genes for MODY1 and MODY3 have not been identified. HNP-4 is a gene mapping within the MODY1 locus
                                              Gaps
                                              .
0
0.4%; Score 15.2; DB 1; Length 20;
85.0%; Pred. No. 2.3e+02;
iive 0; Mismatches 3; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         0.4%; Score 15; DB 1; Length 17; 100.0%; Pred. No. 1.9e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 17 BP; 2 A; 3 C; 8 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                 Nucleotide sequence of the PCR primer 15.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Disclosure; Page 86; 116pp; English.
                                                                                      1857 CAGCATTTTCCAAGTAGTCT 1876
                                                                                                              CAGCACTTCCCAAGTAGCCT 1
                                                                                                                                                                                                                                     AAV36994 standard; DNA; 17 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      96US-00749430.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (MILL-) MILLENNIUM PHARM INC.
                                                                                                                                                                                                                                                                                                                       (first entry)
                                            Matches 17; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Thomas J;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 1998-297964/26.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Best Local Similarity
                        Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           MODY3 gene; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Glucksman AM,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WO9821363-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      15-NOV-1996;
                                                                                                                                                                                                                                                                                                                         24-SEP-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              14-NOV-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    22-MAY-1998.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           encoded by
                                                                                                                                                                                                                                                                               AAV36994;
                                                                                                                             20
    Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
                                                                                                                                                                                             RESULT 3
                                                                                                                                                                                                                                       ò
                                                                                                                             셤
```

ð g

```
The present invention relates to enzymatic and antisense nucleic acid molecules that act as inhibitors of the expression of repressor genes encoding the TR2 Orphan receptor, EAR3/COUP-TF-1, the GATA transcription factor gene, IRP-2 and/or the CAATT Displacement Protein (CDP). Inhibition of the repressors removes prevents inhibition (and consequently increases expression of) genes involved in the production of erythropoietin, granulocyte colony stimulating factor protein and interferon alpha
                                                                                                                                                                                                                                                                                                                        Enzymatic and antisense nucleic acid inhibition of repressor genes, useful for producing e.g. granulocyte colony stimulating factor protein, interferon alpha and erythropoietin.
   Ribozyme; erythropoietin; granulocyte colony stimulating factor;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Ribozyme; erythropoietin; granulocyte colony stimulating factor; interferon alpha; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0.4%; Score 15; DB 1; Length 17; 100.0%; Pred. No. 1.9e+02; ive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 17 BP; 5 A; 4 C; 4 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                          Mcswiggen J;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Zwick M, Pavco P, Mcswiggen J;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Hammerhead ribozyme substrate #1725.
                                                                                                                                                                                                                                                                                                                                                                                          Claim 4; Page 105; 164pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 1896 CAGAATGACTTTGCT 1910
                                                                                                                                                                                                                                                        Pavco P,
                                                                                                                                                      11-APR-2000; 2000WO-US009721.
                                                                                                                                                                                      99US-0129390P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           11-APR-2000; 2000WO-US009721
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AAF04209 standard; DNA; 17
                                                                                                                                                                                                                     (RIBO-) RIBOZYME PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                15 cagaargacrirgcr 1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (RIBO-) RIBOZYME PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Local Similarity 100.
les 15; Conservative
                     interferon alpha; ss
                                                                                                                                                                                                                                                                                        WPI; 2000-647423/62.
                                                                                                                                                                                                                                                       Zwick M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2000-647423/62.
                                                                                  WO200061729-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WO200061729-A2
                                                                                                                                                                                        12-APR-1999;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           12-APR-1999;
                                                                                                                   19-OCT-2000.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       16-FEB-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            19-OCT-2000.
                                                                                                                                                                                                                                                   Blatt L,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Blatt L,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AAF04209
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Best Local
Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AAF04209/c
   ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ô
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            This is the nucleotide sequence of the PCR primer used for amplification in the method of the invention, which involves modulating the expression of hepatic nuclear factor or other diabetes related gene. The method is used to treat early onset type II diabetes and defects in insulin secretion. It is based on the discovery that certain mutations in the HNFI gene, encoding a transcription factor, are involved in these conditions
   Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Treating type II diabetes with agent - useful for, e.g. modulating expression of hepatic nuclear factor or other diabetes-related gene.
                                                                                                                                                                                                                                                                                    PCR; primer, amplification; hepatic nuclear factor; HNF; diabetes; type II diabetes; HNFI gene; transcription factor; insulin; ss.
   ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Score 15; DB 1; Length 17;
Pred. No. 1.9e+02;
0; Mismatches 0; Indels
 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 17 BP; 2 A; 3 C; 8 G; 4 T; 0 U; 0 Other;
Mismatches
                                                                                                                                                                                                                                                     Nucleotide sequence of 3' PCR primer 18.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Hammerhead ribozyme substrate #2173.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Disclosure; Page 80; 113pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0.4%; Scur
100.0%; Pre
0;
 ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       96US-00749431.
96US-00760246.
97US-00782047.
                                                                                                                                                   BP.
                                 416
                                                                                                                                                                                                                                                                                                                                                                                                                                         97WO-US020532
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           96US-00748229
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (MILL-) MILLENNIUM PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       CTGCAGGTGCTGGAT 416
                                                                2 CTGCAGGTGCTGGAT 16
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         CTGCAGGTGCTGGAT 16
                                                                                                                                                 AAV41419 standard; DNA; 17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AAF04657 standard; DNA; 17
                                                                                                                                                                                                                  24-SEP-1998 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            16-FEB-2001 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Local Similarity 100.
nes 15; Conservative
 15; Conservative
                               402 CTGCAGGTGCTGGAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            WPI; 1998-297866/26.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Glucksmann AM;
                                                                                                                                                                                                                                                                                                                                                                     WO9821239-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                        07-NOV-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           12-NOV-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         15-NOV-1996;
04-DEC-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          10-JAN-1997;
                                                                                                                                                                                                                                                                                                                                                                                                        22-MAY-1998.
                                                                                                                                                                                                                                                                                                                                      Synthetic
                                                                                                                                                                                  AAV41419;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       402
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   N
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AAF04657
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      RESULT 336
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Matches
 Matches
```

AAF04657/
ID AAF0
XX
AC AAF0
XX
DT 16-F
XX
XX
XX

ઠે

ö

Gaps

ö

```
The present invention relates to enzymatic and antisense nucleic acid molecules that act as inhibitors of the expression of repressor genes encoding the TR2 Orphan receptor, EAR3/COUP-TF-1, the GATA transcription factor gene, IRF-2 and/or the CAATT Displacement Protein (CDP). Inhibition of the repressors removes prevents inhibition (and consequently increases expression of) genes involved in the production of erythropoletin, granulocyte colony stimulating factor protein and
Enzymatic and antisense nucleic acid inhibition of repressor genes, useful for producing e.g. granulocyte colony stimulating factor protein, interferon alpha and erythropoietin.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Seguence 17 BP; 5 A; 4 C; 4 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                        Claim 4; Page 95; 164pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       interferon alpha
```

Score 15; DB 1; Length 17; Pred. No. 1.9e+02; 0; Mismatches 0; Indels Ouery Match 0.4%; Score 15; DB Best Local Similarity 100.0%; Pred. No. 1.9 1896 CAGAATGACTTTGCT 1910 15 cagaargacrtrgcr 1 원

ô

Gaps

;

ABK03219 standard; RNA; 17 12-MAR-2002 (first entry) Human CD20 Inozyme #170. ABK03219;

ВР

Human; ss; antisense therapy, cytostatic; antiinflammatory; haemostatic; cerebroprotective; anotropic; neuroprotective; antiparkinsonian; muscular; CD20; neurite growth inhibitor gene; NOGO; hammerhead ribozyme; DNAzyme; inozyme; G-cleaver; amberzyme; zinzyme; lymphoma; leukaemia; human; mnunodeficiency virus; HIV associated HHL; lymphoma; leukaemia; human; mnunodeficiency virus; HIV associated HHL; mantle-cell lymphoma; MCL; immunot thrombocytopaenia; stroke; dementia; inflammatory arthropathy; central nervous system injury; celebrotropachic; celebrotropachic; dimentia; disease; multiple sclerosis; chemotherapy-induced neuropathy; amyotrophic lateral sclerosis; ALS; parkingon's disease; ataxia; Huntington's disease; Creutzfeldt-Jakob disease; muscular dystrophy; neurodegenerative disease. 

sapiens. Synthetic.

WO200159103-A2.

16-AUG-2001.

09-FEB-2001; 2001WO-US004273

11-FEB-2000; 2000US-0181797P. 28-FEB-2000; 2000US-0185516P. 06-MAR-2000; 2000US-0187128P

RIBOZYME PHARM INC. CHOWRIRA B M. MCSWIGGEN J. BLATT L. RIBO-) CHOM/) (BLAT/) (MCSM/)

Chowrira BM; Blatt L, Mcswiggen J,

WPI; 2001-607195/69

Nucleic acid molecules, e.g., enzymatic nucleic acids and antisense

The INVENTION FIGURE to a nucleic acid molecule which down regulates expression of a neurite growth inhibitor gene (NOGO). The INVENTION FIGURE acids may be enzymatic nucleic acid molecule which down regulates acids may be enzymatic nucleic acids (e.g. a ribozyme or a nucleic acids is may be enzymatic nucleic acid cleaving an RNA molecule possessing an NCH motif), a G-cleaver (cleaving RNA with a NYN motif) promain an an NCH motif). The CD20-targetting nucleic acid is used to cleave RNA with a YGY motif). The CD20-targetting nucleic acid is used to cleave RNA of CD20 in the presence of a divalent cation that is preferably Mg^2+. The coll and treat a patient having a condition associated with the level of CD20. The treatment may further comprise the use of one or more contragines. In particular, the CD20 targetting nucleic acid may be used to treat lymphoma, leukaemia, B-cell lymphoma, low-grade or follicular non-creat lymphoma (MLL), bulky low-grade or follicular non-creat lymphoma (MLL), immunocytopenia, modeficiency virus) associated NHL, mantle-cell lymphoma (MCL), immunocytopenia, and inflammatory arthropathy. The NOGO-created with a cell to reduce NOGO ace in the presence of a divalent cation that is preferably Mg^2+. Furthermore, the presence of a divalent cation that is preferably Mg^2+. Furthermore, the condition associated with the level of cell and treat a patient having a condition associated with the level of NOGO. The treatment may further comprise the use of one or more cell and treatment may further comprise the use of one or more contented with a cell of acid any be used to cleave NOGO. The treatment may further comprise the use of one or more cell and treatment may further comprise the use of one or more cell and treatment may further comprise the use of one or more contented with a cell of acid any because, and entain and indices and may be used to cleamether may further comprise the use of one or more cell and erropathy, and/or other hardon or solvens and encoded or condition of condition aci constructs, which down regulate expression of a CD20 gene or neurite growth inhibitor gene useful for treating, e.g., lymphoma, leukemia, and central nervous system injury. The invention relates to a nucleic acid molecule which down regulates Sequence 17 BP; 3 A; 5 C; 3 G; 0 T; 6 U; 0 Other; sequence is an inozyme of the invention Claim 30; Page 148; 200pp; English 

Gaps ; 0 0.4%; Score 15; DB 1; Length 17; 00.0%; Pred. No. 1.9e+02; 0; Indels Query Match
Best Local Similarity 100.0%; Fred. No. 1.9

ö

3797 CTGACAGGAGAACTA 3811 17 CTGACAGGAGAACTA 3 ò g ABK03220 standard; RNA; 17 BP 12-MAR-2002 (first entry)

Human CD20 Inozyme #171.

Human; ss; antisense therapy; cytostatic; antiinflammatory; haemostatic; cerebroprotective; antiparkinsonian; muscular; CD20; neurite growth inhibitor gene; NOG0; hammerhead ribozyme; DNAzyme; inozyme; G-cleaver; amberzyme; zinzyme; lumphoma; leukaemia; buckayme; inozyme; deaver; amberzyme; zinzyme; lymphoma; leukaemia; human; immunodeficiency virus; HIV associated NHL; lymphocytic leukaemia; MCL; immunocytoma; MC; immune thrombocytopaenia; stroke; dementia; inflammatory arthropathy; central nervous system injury; central nervous system injury; characylanduced neuropathy; amplementy af disease; multiple sclerosis; chemocherapy-induced neuropathy; amplementy adisease; ataxia; Huntington's disease; 

Creutzfeldt-Jakob disease; muscular dystrophy; neurodegenerative disease.

Homo sapiens

```
11-FEB-2000; 2000US-0181797P.
28-FEB-2000; 2000US-0185516P.
06-MAR-2000; 2000US-0187128P.
                     09-FEB-2001; 2001WO-US004273
                                           (RIBO-) RIBOZYME PHARM INC.
                                                               Blatt L, Mcswiggen J,
                                                       (CHOW/) CHOWRIRA B M.
                                                   MCSWIGGEN J.
                                                                      WPI; 2001-607195/69.
                                               BLATT L.
       WO200159103-A2.
              16-AUG-2001.
Synthetic.
                                                BLAT/)
                                                    MCSW/)
```

```
ö
                                        Gaps
                                        ;
0
0.4%; Score 15; DB 1; Length 17;
100.0%; Pred. No. 1.9e+02;
tive 0; Mismatches 0; Indels
                    Best Local Similarity 100. Matches 15; Conservative
     Query Match
```

The invention relates to a nucleic acid molecule which down regulates expression of a CD20 gene and a nucleic acid molecule which down regulates expression of a neurite growth inhibitor gene (NGO2). The regulates expression of a neurite growth inhibitor gene (NGO2). The nucleic acids may be enzymatic nucleic acids (e.g. a ribozyme or a DNAzyme) an Inozyme (an endolytic nucleic acid cleaving an RNA molecule possessing an NCH motif), a G-cleaver (cleaving RNA with a NRY motif) propersion of CD20 in the presence of a divalent cation that is preferably MG 2+.

The coll and treat a patient having a condition associated with the level of CD20. The treatment may further comprise the use of one or more cherapies. In particular, the CD20 targetting nucleic acid may be used to treat lymphoma (NHL), bulky low-grade or follicular NHL, lymphocytic treat lymphoma (NHL), bulky low-grade or follicular NHL, lymphocytic lymphoma (NHL), immunocytoma (IMC), small B-cell lymphocytic lymphoma, comparing an anticle acid may be contacted with a cell to reduce NGO gene in the presence of a divalent cation that is preferably MG 2+. Furthermore, the crucial may be contacted with a cell to reduce NGO gene in the presence of a divalent cation that is preferably MG 2+. Furthermore, the coulcid may be contacted with a condition associated with the level of NOGO. The treatment may further comprise the use of one or more cell and treat a patient having a condition associated with the level of NOGO. The treatment may further comprise the use of one or more coll and treat a patient having a condition associated with the level of NOGO. The treatment may further comprise the use of one or more coll and treat a patient having a condition associated with the level of theraples. In particular, the NOGO-targetting nucleic acid may be used to treat central nervous system (CNA) injury and cerebrovascular accident cation the modular dystem (CNA) alsease, ataxia, Huntington's disease, central encodes ataxia, Huntington's disease, central encodes ataxia, H Nucleic acid molecules, e.g., enzymatic nucleic acids and antisense constructs, which down regulate expression of a CD20 gene or neurite growth inhibitor gene useful for treating, e.g., lymphoma, leukemia, and sequence is an inozyme of the invention Claim 30; Page 148; 200pp; English. central nervous system injury.

Sequence 17 BP; 3 A; 5 C; 3 G; 0 T; 6 U; 0 Other;

3797 CTGACAGGAGAACTA 3811 CTGACAGGAGAACTA 1

15

ò 셤 RESULT 340 ABK02853/c

ABK02853 standard; RNA; 17 BP

ABK02853;

(first entry) 12-MAR-2002 Human CD20 Hammerhead ribozyme #152.

Human; ss; antisense therapy; cytostatic; antiinflammatory; haemostatic; cerebroprotective; notropic; neuroprotective; antiparkinsonian; muscular; CD20; neurite growth inhibitor gene; NOGO; hammerhead ribozyme; DNAzyme; inczyme; inczyme; inczyme; loukaemia; B-cell lymphoma; non-Hodgkin's lymphoma; NHL; lymphoma; leukaemia; B-cell lymphoma; non-Hodgkin's lymphoma; NHL; lymphocytic leukaemia; MCL; immunocytcoma; IMC; immune thrombocytcopaemia; stroke; dementia; inflammatory arthropathy; central nervous system injury; cerebrovascular accident; CVA; Alzheimer's disease; multiple sclerosis; chemotherapy-induced neuropathy; amyotrophic lateral sclerosis; ALS; Parkinson's disease; ataxia; Huntington's disease; creutzfeldt-Jakob disease; muscular dystrophy; neurodegenerative disease.

Homo sapiens Synthetic.

Chowrira BM;

WO200159103-A2.

16-AUG-2001.

19-FEB-2001; 2001WO-US004273.

11-FEB-2000; 2000US-0181797P. 28-FEB-2000; 2000US-0185516P. 06-MAR-2000; 2000US-0187128P.

RIBOZYME PHARM INC. BLATT L. (RIBO-)

BLAT/)

(MCSW/) MCSWIGGEN J. (CHOW/) CHOWRIRA B M.

Chowrira BM; Blatt L, Mcswiggen J,

WPI; 2001-607195/69

Nucleic acid molecules, e.g., enzymatic nucleic acids and antisense constructs, which down regulate expression of a CD20 gene or neurite growth inhibitor gene useful for treating, e.g., lymphoma, leukemia, and central nervous system injury.

Claim 30; Page 142; 200pp; English.

The invention relates to a nucleic acid molecule which down regulates expression of a CD20 gene and a nucleic acid molecule which down cegulates expression of a neurite growth inhibitor gene (NGCD). The cegulates expression of a neurite growth inhibitor gene (NGCD). The concleic acids may be enzymatic nucleic acids (e.g. a ribozyme or a DNzyme) an Inozyme (an endolytic nucleic acid cleaving an RNA molecule possessing an NCH motif), a G-cleaver (cleaving RNA with a NCM motif). The CD20-targetting nucleic acid is used to cleave RNA with a YCY motif). The CD20-targetting nucleic acid is used to cleave RNA of CD20 in the presence of a divalent cation that is preferably Mg^2+. The CD20-targetting nucleic acid is used to cleave RNA of CD20. The treatment may further comprise the use of one or more the cell and treat a patient having a condition associated with the level of CD20. The treatment may further comprise the use of one or more therefore in particular, the CD20 targetting nucleic acid may be used to treat lymphoma, leukaemia, B-cell lymphoma, low-grade or follicular NHL, lymphocytic leukaemia, HIV (human immunodeficiency virus) associated NHL, mantle-cell lymphoma (NHL), immunocytoma (IMC), small B-cell lymphocytic lymphoma, cimmune thrombocytopaenia, and inflammatory archropathy. The NOGO gene in the concent of a divalent cation that is preferably Mg^2+. Furthermore, the nucleic acid may be contacted with a collicular of nucleic acid may be contacted with a condition associated with the level of cell and treat a patient having a condition associated vict the level of noce

```
oligonucleotides can also be used to quantify raf expression in assays
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    WPI; 2002-749291/81.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Watt AT;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  US6444466-B1
                                                                                                                                                                                                                                                                                                                                                                                                                                                        13-FEB-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   03-SEP-2002
                                                                                                                                                                                                                                                                                                                                                                                                          ABT13950;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        4
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Ward DT,
                                                                                                      Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match
                                                                                                                               Best Loca
Matches
                                                                                                                                                                                                                                                                                                               RESULT 342
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 343
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Matches
                                                                                                                                                                                                                                                                                                                                          ABT13950
         ប្ដង្គន្ល
                                                                                                                                                                                                                                        요
                                                                                                                                                                                              à
                                                                                                                                                                                                                                                                                                                                                                à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        셤
                                                                                                                                                                                                                                                                                                               ö
therapies. In particular, the NOGO-targetting nucleic acid may be used to treat central nervous system (CNS) injury and cerebrovascular accident (CVA, stroke), Alzheimer's disease, dementia, multiple sclerosis (MS), chemotherapy-induced neuropathy, amyotrophic lateral sclerosis (ALS), Parkinson's disease, ataxia, Huntington's disease, Creutzfeldt-Jakob disease, muscular dystrophy, and/or other neurodegenerative disease states which respond to the modulation of NOGO expression. The present
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         This sequence represents an example of an antieense oligomucleotide of the invention. The oligomucleotides are 8-50 nucleotides in length, and are targeted to a nucleic acid encoding human B-raf and which is capable of inhibiting human B-raf expression. The oligomucleotides is used to inhibit the (abnormal) expression of human B-raf, to inhibit hyperproliferation of cells, to treat or prevent an abnormal proliferation of cells, to treat or prevent an abnormal proliferation condition, e.g. hyperproliferative disorders such as cancer (e.g. of the brain or nervous system), restenosis, psoriasis or a disorder characterised by T-cell activation and growth. They may also be used to diagnose these diseases, as well as atherosclerosis. The oligomucleotides of the invention may be used to distinguish raf-associated tumours from tumours having other etiologies. The antisense
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Antisense oligonucleotide, B-raf, human; inhibitor; T-cell activation;
                                                                                                                                                                                                                                                                                                               Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             antisense oligonucleotides - for modulation of human B-raf gene
                                                                                                                                                                                                                                                                                                             .
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        hyperproliferative disorder; cancer; restenosis; psoriasis; atherosclerosis; taf-associated tumour; diagnosis; therapy; ss.
                                                                                                                                                                                                                                                                Length 17;
                                                                                                                                                                                                                                                                                                        0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human B-raf kinase antisense oligonucleotide Isis#14141.
                                                                                                                                                                   sequence is a hammerhead ribozyme of the invention
                                                                                                                                                                                                                Sequence 17 BP; 3 A; 5 C; 3 G; 0 T; 6 U; 0 Other;
                                                                                                                                                                                                                                                              Score 15; DB 1; Lo
Pred. No. 1.9e+02;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  a
"phosphorothioate bases"
                                                                                                                                                                                                                                                            Query Match 0.4%; Score 15; DB Best Local Similarity 100.0%; Pred. No. 1.9 Matches 15; Conservative 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Disclosure; Page 22; 72pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          BP.
                                                                                                                                                                                                                                                                                                                                                      3797 CTGACAGGAGAACTA 3811
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              98WO-US013961
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        97US-00888982.
                                                                                                                                                                                                                                                                                                                                                                                                     16 crcacacadaacra 2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        AAX21944 standard; DNA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /note=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 WPI; 1999-120502/10.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Key
modified_base
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     18-MAY-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         06-JUL-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     07-JUL-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WO9902167-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             21-JAN-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Synthetic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        expression
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AAX21944;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Monia BP;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              RESULT 341
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAX21944/C

ID AAX22

AC AAX22

XX A
    SSSSSSSSSS
                                                                                                                                                                                                                                                                                                                                                                                                  셤
                                                                                                                                                                                                                                                                                                                                                      ò
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      The invention comprises antisense oligonucleotides which are targeted to the coding region of the human helicase-moi gene. The antisense oligonucleotides of the invention are useful for inhibiting the expression of human helicase-moi in cells or tissues, and for traating a helicase-moi-associated condition. The antisense oligonucleotides of the invention may also be used to delay infection, inflammation and tumour formation. The present DNA sequence represents a human helicase-moi gene antisense oligonucleotide of the invention. NOTE: The present DNA sequence has a phosphorothicate backbone, bases 1-5 and 16-20 are 2'-methoxyethyl (2'-MOE) nucleotides
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Novel antisense compound for modulating expression of human helicase-moi and for treating inflammation, specifically hybridizes to a specific region in nucleic acid molecule encoding the human helicase-moi.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Human; antisense gene therapy; phosphorothioate backbone; antisense oligonucleotide; helicase-moi gene; inflammation; ss; helicase-moi-associated condition; infection; tumour formation; 2-MOE nucleotide; 2'-methoxyethyl nucleotide.
                                                                                                                             ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ó;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  0.4%; Score 15; DB 1; Length 20; 100.0%; Pred. No. 2.4e+02; ive 0; Mismatches 0; Indels
                                                             Length 20;
                                                                                                                          0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human helicase-moi inhibiting oligonucleotide #75.
Sequence 20 BP; 3 A; 6 C; 1 G; 10 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 20 BP; 6 A; 5 C; 3 G; 6 T; 0 U; 0 Other;
                                                             0.4%; Score 15; DB 1; Le
.00.0%; Pred. No. 2.4e+02;
.ve 0; Mismatches 0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Claim 3; Col 45-46; 52pp; English.
                                                                0.1.,
100.0%; Pre-
                                                                                                                                                                                                                                                                                                                                                                                                                           ВР
                                                                                                                                                                                              1467 AAACAAATGAGTGAG 1481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          2331 ATCCATGAAGGTTTC 2345
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     10-MAY-2001; 2001US-00853768.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      10-MAY-2001; 2001US-00853768.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Arccardaderric 18
                                                                                                                                                                                                                                                                                                                                                                                                                        ABT13950 standard; DNA; 20
                                                                                                                                                                                                                                                        20 AAACAAATGAGTGAG 6
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      (first entry)
                                                                                           Local Similarity 100.
1es 15; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      15; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (ISIS-) ISIS PHARM INC
```

```
Preventing and/or treating conditions associated with raf expression, such as hyperproliferative disorders, atherosclerosis and tumors, using antisense oligonucleotide modulation of human raf gene expression.
                                                                                              tumour metastasis, human; raf; raf expression inhibitor; cytostatic; antiarteriosclerotic; antisense-therapy; hyperproliferative disorder; atherosclerosis; tumour; b-raf kinase; antisense oligonucleotide; ss.
                                                     Human b-raf kinase antisense oligonucleotide seq id 66.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Example 18; SEQ ID NO 93; 41pp; English
                                                                                                                                                                                                                                                                                                                                  14-JUN-2002; 2002US-00173225.
    12-FEB-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 2003-863446/80
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (MONI/) MONIA B P.
                                                                                                                                                                                                                                           US2003119769-A1.
                                                                                                                                                                                               Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                             31-MAY-1994;
31-MAY-1995;
26-NOV-1996;
07-JUL-1997;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             25-JAN-2002;
                                                                                                                                                                                                                                                                                      26-JUN-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            06-JUL-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               28-AUG-1998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Monia BP;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Matches
         NAMES OF COLORS OF STREET STRE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        are targetted to nucleic acids encoding human raf proteins and capable of inhibiting raf expression. The invention also relates to methods of inhibiting hyperproliferation of cells which involves contacting the hyperproliferating cells with a therapeutically effective amount of an oligonuclectide of the invention. The method is useful for treating cancer, angiogenesis or neovascularisation, especially ocular angiogenesis or neovascularisation. The present DNA sequence is an antisense oligonuclectide targetted to human B-raf kinase
                                                                                                                                                                                                           Human; raf; hyperproliferation; neovascularisation; ocular angiogenesis; therapy; cancer; cytostatic; anti-angiogenic; vascular; ophthalmological; antisense; phosphorothioate backbone; B-raf kinase; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         present invention relates to novel antisense oligonucleotides which
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Treating cancer, angiogenesis or neovascularization by administering antisense oligonucleotides targeted to human raf sequences.
                                                                                                                                                                  Human B-raf kinase antisense oligonucleotide ISIS #14141.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.4%; Score 15; DB 1; Length 20;
100.0%; Pred. No. 2.4e+02;
ve 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                 /mod_base= OTHER
/note= "Phosphorothioate backbone"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 20 BP; 3 A; 6 C; 1 G; 10 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                             Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Example 18; Col 26; 41pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    0.7.,
100.0%; Pre-
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                95WO-US007111.
96US-00756806.
97US-00888982.
98WO-US013961.
98US-00143214.
                          AAD44803 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               18-FEB-2000; 2000US-00506073
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             94US-00250856
                                                                                                                      (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                              đ
                                                                                                                                                                                                                                                                                                                                                                                                      . 20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WPI; 2002-597918/64.
                                                                                                                                                                                                                                                                                                                                                                             Key
modified_base
                                                                                                                    13-DEC-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     31-MAY-1995;
26-NOV-1996;
07-JUL-1997;
                                                                                                                                                                                                                                                                                                          Homo sapiens.
Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     US6410518-B1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             31-MAY-1994;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 25-JUN-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 28-AUG-1998;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         06-JUL-1998
                                                                        AAD44803;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Monia BP;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Query Match
AAD44803/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         The
```

94US-00250856. 95WO-US007111. 96US-00756806. 97US-00888982. 98WO-US013961.

2000US-00506073 2002US-00057550

```
ö
The invention describes a method of preventing or treating tumour metastasis in an animal comprising administering to the animal an oligomucleotide 8-50 nucleotides in length, which is targeted to mRNA encoding human raf and capable of inhibiting raf expression. Also disclosed are raf oligonucleotides, nucleic acids, proteins and compositions used in the methods of the invention. The oligonucleotides have cytostatic and antiarteriosclerotic properties, are useful as Rafinhibitors and in antisense-therapy. The methods and compositions of the present invention are useful for preventing and/or treating conditions associated with raf expression, such as hyperproliferative disorders, atherosclerosis and tumourer. This sequence represents a human b-raf
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human, antisense, lung dysfunction; nasal airway dysfunction; antiinflammatory steroid; ubiquinone; antiinflammatory; antiallergic;
                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                                                            Length 20;
                                                                                                                                                                                                                                                                                                                                                                                                                                           0; Indels
                                                                                                                                                                                                                                                                                                                                          Sequence 20 BP; 3 A; 6 C; 1 G; 10 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                      Score 15; DB 1; Le
Pred. No. 2.4e+02;
0; Mismatches 0;
                                                                                                                                                                                                                                                                                                                                                                       0.4%; Scor
100.0%; Pred
0; F
                                                                                                                                                                                                                                                                                           kinase antisense oligonucleotide.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human oligonucleotide sequence.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ABZ86360 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1467 AAACAAATGAGTGAG 1481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                             Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              20 AAACAAATGAGTGAG
                                                                                                                                                                                                                                                                                                                                                                                                           Local Similarity
tes 15; Conserva
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             17-OCT-2003
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ABZ86360;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 345
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ABZ86360/c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          MAKEK BK BK KK ID
```

ö

Gaps

ö

0; Indels

1467 AAACAAATGAGTGAG 1481

Conservative

Local Similarity

15;

Best Loc Matches

AAACAAATGAGTGAG 6

20

g ð

ADF09797 standard; DNA; 20 BP

ADF09797

RESULT 344
ADF09797/C
ID ADF097
XX
AC ADF097
XX

**新**罗斯斯克

10001863-3.sl.rng

```
The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation coodon, coding regalon, 5' or 3' end genomic flanking regions, 5' and 3' intron-exon junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or nasal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallergic, antiathmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a use in antisense gene therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of an exceptor, producing sensitivity to adenosine, reducing levels of adenosine receptor, producing sensitivity to adenosine, reducing levels of adenosine receptor, producing bronchoodilation, increasing levels of ubiquinone or lung sunfactant in a subject, for reducing levels of adenosine receptor, producing sensitivity to adenosine, reducing levels of adenosine receptor, producing sensitivity to adenosine, reducing levels of adenosine receptor, producing allergies, or a respiratory disease or condition, lung inflammation, but was obtained in electronic format directly from WIPO
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or
antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy;
                                                                                                                                                                                                                                                                                                                                                                                                    Aguilar D;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         0.4%; Score 15; DB 1; Length 20; 100.0%; Pred. No. 2.4e+02; ive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                    Katz E, Pabalan J,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Sequence 20 BP; 5 A; 7 C; 6 G; 2 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ftp.wipo.int/pub/published_pct_sequences
                                                                      lung inflammation; respiratory disease; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Claim 15; SEQ ID NO 1602; 872pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                          ŝ
                                                                                                                                                                                                                                                                                                                                                                                                 Sandrasagra A,
L, Shahabuddin
                                                                                                                                                                                                                                                            23-APR-2002; 2002WO-US013135.
                                                                                                                                                                                                                                                                                                        24-APR-2001; 2001US-0286137P.
                                                                                                                                                                                                                                                                                                                                                      (EPIG-) EPIGENESIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Local Similarity 100.
ses 15; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2003-229219/22.
                                                                                                                                                                                                                                                                                                                                                                                                    Li Y, Sa
Tang L,
                                                                                                                                                                WO200285308-A2
                                                                                                                                                                                                              31-OCT-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ubiquinone.
                                                                                                                                                                                                                                                                                                                                                                                                                          Miller S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
                                                                                                                                                                                                                                                                                                                                                                                                    Nyce JW,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Best Loca
Matches
```

```
        Query Match
        0.4%; Score 15; DB 1; Length 20;

        Best Local Similarity 100.0%; Pred. No. 2.4e+02;

        Matches 15; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

        Qy 2883 GTCTTCCAGGTGGC 2897

        Db 20 GTCTTCCAGGTGGC 6

        RESULT 346

        ABZ86408/C

        ID ABZ86408

        XX

        AC ABZ86408; XX

        XX

        DE Human oligonucleotide sequence.

        XX

        Y

        XX

        XX
```

```
The invention relates to a novel pharmaceutical composition, which has a first active agent comprising an oligonucleotide antisense to the initiation codon, coding region, 5' or 3' end genomic flanking regions, 5' and 3' end 3' intron-excn junctions, or regions within 2-10 nucleotides of junctions of genes encoding a polypeptide associated with lung and/or masal airway dysfunction and a second active agent comprising an antiinflammatory steroid and ubiquinone. A composition of the invention has antiinflammatory, antiallargic, antiasthmatic, hypotensive, immunosuppressive, and cytostatic activity. The composition may have a use in antisense gene therapy. The composition is useful for treating or preventing a respiratory, lung or malignant disease or condition, also for enhancing the prophylactic or therapeutic respiratory effect of an artifinammatory steroid in a subject, for reducing of depleting levels of or reducing sensitivity to adenosine, reducing levels of adenosine receptor, producing bronchodilation, increasing levels of ubiquinone or lung surfactant in a subject, a tissue, or treating bronchoconstriction, lung allergies, or a respiratory disease or condition. Note: The sequence data for this parent is not represented in the printed specification, but was obtained in electronic format directly from WIPO
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Pharmaceutical composition for treating ailments associated with impaired respiration, has oligo(s) antisense to specific gene(s) or its corresponding RNAs, and glucocorticoid or non-glucocorticoid steroid or
antiasthmatic; hypotensive; immunosuppressive; cytostatic; gene therapy; antisense gene therapy; respiratory; lung; adenosine sensitivity; adenosine receptor; bronchodilation; bronchoconstriction; lung allergy; lung inflammation; respiratory disease; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human, 88; antisense; c-raf; a-raf; b-raf; protein kinase; cancer; signal transduction; cell proliferation; lung carcinoma; cytostatic;
                                                                                                                                                                                                                                                                                                                                                                     Katz E, Pabalan J, Aguilar D;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           .
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Antisense oligonucleotide targeting human b-raf, ISIS14141.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15; DB 1; Length 20; 100.0%; Pred. No. 2.4e+02; ive 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Sequence 20 BP; 13 A; 2 C; 1 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ftp.wipo.int/pub/published_pct_sequences
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Claim 15; SEQ ID NO 1650; 872pp; English.
                                                                                                                                                                                                                                                                                                                                                                   Li Y, Sandrasagra A, Ka
Tang L, Shahabuddin S;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ACD42186 standard; DNA; 20 BP.
                                                                                                                                                                                                                                        23-APR-2002; 2002WO-US013135.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    3305 TTTTTATTTTATAT 3319
                                                                                                                                                                                                                                                                                   24-APR-2001; 2001US-0286137P.
                                                                                                                                                                                                                                                                                                                           (EPIG-) EPIGENESIS PHARM INC.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         15; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              15 TTTTATTTTATAT
                                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 2003-229219/22.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Best Local Similarity
                                                                                                                                                     WO200285308-A2
                                                                                                             Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     05-SEP-2003
                                                                                                                                                                                              31-OCT-2002
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ubiquinone.
                                                                                                                                                                                                                                                                                                                                                                                       Miller S,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACD42186;
                                                                                                                                                                                                                                                                                                                                                                     Nyce JW,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        RESULT 347
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ACD42186/
    à
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   MAKAKAKAKA
MAKAKAKA
MAKAKA
MAKA
```

```
Human helicase-moi, antisense oligonucleotide #75
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          RESULT 349
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Matches
  ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 셤
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     which is targeted to mRNA encoding human craft, a-raf or b-raf (raf is a protein kinase playing a regulatory role in signal transduction, regulating cell proliferation and has been implicated in lung carcinoms), and which is capable of inhibiting raf expression. Also included is a composition comprising the oligonuclectide and a pharmaceutically acceptable carrier. The antisense oligonuclectide is useful for this inhibiting the expression of human raf in human cells or tissues, by contacting the human cells or tissues with the oligo. The oligo. is also is useful for treating or preventing a disease or condition associated with the expression of raf by administering it in combination with a chemotherapeutic agent to a human or cells of the human, where the expression of raf by administering of the condition is a hyperproliferative condition such as cancer, angiogenesis or neovascularisation (preferably ocular angiogenesis or neovascularisation). The oligo. is also useful for inhibiting hyperproliferation of cells. The oligos are also useful as tools, for example for detecting and determining the role of raf expression in various cell functions and physiological processes and conditions and for determining the role of raf expression in the condition and physiological processes and conditions and for determining the role of each example.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                 Novel antisense oligonucleotide which is targeted to mRNA encoding human raf and which is capable of inhibiting raf expression, useful for treating or preventing hyperproliferative conditions such as cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  diagnosing conditions associated with raf expression and for research purposes. The present sequence is an antisense oligonucleotide targeting
                                                                                                                                                                                                                                                                                                                                                                                                             invention relates to an oligonucleotide 8-50 nucleotides in length
antisense gene therapy, chemotherapeutic agent, angiogenesis, hyperproliferative condition, neovascularisation, ocular angiogenesis
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Score 15; DB 1; Length 20;
Pred. No. 2.4e+02;
0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Sequence 20 BP; 3 A; 6 C; 1 G; 10 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                     Example 18; Page 14; 42pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               0.4°,
100.0%; Pr
                                                                                                                                                        95WO-US007111.
96US-00756806.
97US-00888982.
98WO-US013961.
98US-00143214.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1467 AAACAAATGAGTGAG 1481
                                                                                                                   2002US-00057550
                                                                                                                                             94US-00250856
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 01-JUL-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  AAACAAATGAGTGAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         15; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ADN60155 standard; DNA;
                                                                                                                                                                                                                                                                                                       WPI; 2003-503332/47.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Local Similarity
                                                                                                                                                                                                                                                    (MONI/) MONIA B P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               a human raf mRNA
                                                                US2003032607-A1
                                                                                                                                           31-MAY-1994;
31-MAY-1995;
26-NOV-1996;
                                                                                                                   25-JAN-2002;
                                       Homo sapiens
                                                                                                                                                                                                                          18-FEB-2000;
                                                                                                                                                                                     07-JUL-1997;
                                                                                                                                                                                              06-JUL-1998;
                                                                                         13-FEB-2003
                                                                                                                                                                                                                                                                           Monia BP;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ADN60155;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  RESULT 348
ADN60155
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Matches
ઠે
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          g
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            BXXXEX
```

```
ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      The invention relates to antisense oligonuclectides, compositions and methods for modulating the expression of helicase-moi. The oligonucleotides are used in treating an animal having a disease or condition associated with hicase-moi, such as inflammation, a hyperproliferative disorder or a condition that arises from RNA-mediated
                                                                                                                                                                                /mod_base= Other
/note= "Phosphorothioate linkages. All cytidines are 5'-
methylcytidines"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              New antisense oligonucleotides for modulating helicase-moi expression, useful for diagnosing, preventing or treating diseases or conditions associated with helicase-moi, e.g. inflammation or hyperproliferative
Cytostatic; Antisense therapy; ss; human; helicase-moi; inflammation; hyperproliferative disorder; RNA-mediated interference; probe.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     interference. The present sequence represents a human helicase-moi
antisense oligonucleotide.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ö
                                                                                                                                                                                                                                                                                             /*tag= a
/mod_base= Other
/note= "2'-methoxyethyl (2'-MOE) nucleotides"
16. 20
                                                                                                                                                                                                                                                                                                                                                                                                                          /mod_base= Other
/note= "2'-methoxyethyl (2'-MOE) nucleotides"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0.4%; Score 15; DB 1; Length 20;
00.0%; Pred. No. 2.4e+02;
ve 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 20 BP; 6 A; 5 C; 3 G; 6 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Human cathepsin C-derived oligo SEQ ID 1602
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Claim 3; SEQ ID NO 88; 56pp; English.
                                                                                                                         Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     100.0%; FIU
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         ABD22590/c
ID ABD22590 standard; DNA; 20 BP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               2331 ATCCATGAAGGTTTC 2345
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       31-MAY-2002; 2002US-00160632.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      .0-MAY-2001; 2001US-00853768.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Local Similarity 100.
16s 15; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               4 ATCCATGAAGGTTTC
                                                                                                                                                   1. .20
/*tag=
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               WPI; 2003-898586/82
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Ward DT, Watt AT;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (WARD/) WARD D T.
(WATT/) WATT A T.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     US2003176380-A1
                                                                                                                         Key
modified_base
                                                                                                                                                                                                                                                                             modified base
                                                                                                                                                                                                                                                                                                                                                                        modified_base
                                                                           Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           29-JUL-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              disorders
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ABD22590,
```

analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; pulmonary transplantation rejection; ss; primer. Human, antisense, bronchoconstriction; allergy, hyposecretion, pain, respiratory tract inflammation, adenosine sensitivity, lung, cancer, surfactant depletion, antiallergic, antiinflammatory, antiasthmatic, 

Homo sapiens.

WO200285309-A2

31-OCT-2002.

23-APR-2002; 2002WO-US013143

24-APR-2001; 2001US-0286036P.

(EPIG-) EPIGENESIS PHARM INC

Katz E, Pabalan J, Aguilar D; ŝ Li Y, Sandrasagra A, Tang L, Shahabuddin Nyce JW, Miller S,

WPI; 2003-093058/08.

Pharmaceutical composition for treating asthma, has antisense oligonucleotide containing less percentage of adenosine, targeted to nucleic acids associated with lung airway or lung dysfunction, and bronchodilating agent.

Claim 15; SEQ ID NO 1602; 763pp; English.

The interior describes a nove composition (a) a liret active agent, comprising oligomuclectides, effective for alleviating control describes, effective for alleviating bronchoconstriction, respiratory tract inflammation, allergies and reducing adenosine (son for a proper control, when administered to a mammal. The oligomuclectides are derived from a gene encoding or regulating control or target polypeptide associated with lung airway or lung dysfunction or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery or cancer and can be anti-sense to the kit. The composition of analgesic, hypotensive, insuppressive and for use of the kit. The composition of instructions for adding a carrier and for use of the kit. The composition of analgesic, hypotensive, immunosuppressive and cytostatic activity, is a confinent or an analgesic, hypotensive, immunosuppressive and cytostatic activity, is a confinent or accomprise oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypeptide present in the lungs. The pulmonary obstruction, and/or bronchoconstriction and/or lung inflammation, allergies and/or surfactant hypoproduction are associated with a disease or condition such as pulmonary vasoconstriction, confinent or specification, pulmonary impeded respiration, respiratory distress syndrome, pain, cystic fibrosis, allergic rhintis, pulmonary contransplantation rejection, pulmonary impeded respiration, respiratory contransplantation rejection, pulmonary impeded respiration, respiratory contransplantation rejection, pulmonary infections, bulmonary disease, pulmonary intercibles by an an or condition such as pulmonary disease, pulmonary contresponding to the anti-sense oligos corresponding to the oligonucleotides into products that free adenosine into the target RNA serves to prevent the breakdown of the oligonucleotides into product and the control of the contro This invention describes a novel composition (a) a first active agent,

Sequence 20 BP; 5 A; 7 C; 6 G; 2 T; 0 U; 0 Other;

Gaps ; Length 20; Indels ö Score 15; DB 1; Le Pred. No. 2.4e+02; 100.0%; Pred. No. 4.7 0.48; 15; Conservative Query Match Best Local Similarity Matches

ö

셤

ABD22638 standard; DNA; 20 BP. RESULT 350 ABD22638/

ABD22638;

(first entry) 29-JUL-2004 Human myosin X-derived oligonucleotide SEQ ID 1650.

analgesic; hypotensive; immunosuppressive; cytostatic; cystic fibrosis; beta-adrenergic agonist; respiratory disease; pulmonary vasoconstriction; respiratory distress syndrome; allergic rhinitis; pulmonary hypertension; emphysema; chronic obstructive pulmonary disease; cancer; bronchitis; Human, antisense, bronchoconstriction, allergy, hyposecretion, pain, respiratory tract inflammation, adenosine sensitivity, lung, cancer, surfactant depletion, antiallergic, antiinflammatory, antiasthmatic; pulmonary transplantation rejection; ss; primer.

Homo sapiens.

WO200285309-A2

31-OCT-2002.

23-APR-2002; 2002WO-US013143.

(EPIG-) EPIGENESIS PHARM INC.

24-APR-2001; 2001US-0286036P.

Sandrasagra A, Katz E, Pabalan J, Aguilar D; , Shahabuddin  $S_i$ Li Y, Sar Tang L, Miller S, Nyce JW,

WPI; 2003-093058/08.

t oligonucleotide containing less percentage of adenosine, targeted ( nucleic acids associated with lung airway or lung dysfunction, and Pharmaceutical composition for treating asthma, has antisense bronchodilating agent

Claim 15; SEQ ID NO 1650; 763pp; English.

This interior describes a novel composition, alleviating describes and bronchoconstriction, respiratory tract inflammation, allergies and bronchoconstriction, respiratory tract inflammation, allergies and bronchoconstriction or hyposecretion, when administered to a mammal. The cutacing adenosine sensitivity, levels of adenosine (A) or (A) or (A) receptors, surfactant depletion or hyposecretion, when administered to a mammal. The oligonucleotides are derived from a gene encoding or regulating of a target polypeptide associated with lung airway or lung dysfunction or cancer and can be anti-sense to the corresponding mRNA. The invention also describes a kit, that comprises: (a) a delivery of instructions for adding a carrier and for use of the kit. The composition of the invention has antiallergic, antiinflammatory, antiasthmatic, of the invention as a national lergic, antiinflammatory, antiasthmatic, beta-adrenergic agonist. The composition is useful for preventing or treating a respiratory, lung or mailgnant disease. The administered composition comprises oligo and is administered to reduce the production or availability, or to increase the degradation of the target mRNA or to reduce the amount of target polypeptide present in the lungs. The pulmonary obstruction, and/or bronchoconstriction and/or lung inflammation, allergies and/or surfactant hypoproduction are associated with a disease or condition such as pulmonary vasoconstriction, finflammation, allergies, asthma, impeded respiration, respiratory distress syndrome, pain, cystic fibrosis, allergic rhinitis, pulmonary hypertension, emphysema, chronic obstructive pulmonary disease, pulmonary transplantation rejection, pulmonary infections, bronchitis or cancer. The reduced adenosine content of the anti-sense oligos corresponding to thymidines present in the target RNA serves to prevent the breakdown of the oligonucleotides into products that free adenosine into the system This invention describes a novel composition (a) a first active agent,

```
Query Match 0.4%; Score 15; DB 1; Length 20;
```

一年 日本の

```
The invention relates to an antisense oligonucleotide targeted to a nucleic acid encoding the human DEXRASI polypeptide which specifically hybridises with the nucleic acid molecule and inhibits the expression of DEXRASI. The antisense oligonucleotide comprises at least one modified internucleoside linkage, i.e. a phosphorothioate linkage, at least one modified sugar moiety, preferably a 2'-0-methoxyethyl sugar moiety, or at least one modified nucleobase compounds are useful for inhibiting the antisense oligonucleotides and compounds are useful for inhibiting the expression of DEXRASI and for treating diseases or conditions associated with expression of DEXRASI. e.g. hyperproliferative disorders such as cancer, neurological disorders or a disease or condition arising from aberrant nitric oxide signalling. The antisense compounds are also useful complylactic applications, e.g. to prevent or delay infection, inflammation or tumour formation. This sequence represents an antisense oligonucleotide target region of the invention.
                                                                                                                                  ö
 tissue environment and thereby, to
                                                                                                                                                                                                                                                                                                                                                                                                                                                          Human; DEXRAS1; 88; antisense oligonucleotide; phosphorothioate linkage;
2'-O-methoxyethyl sugar moiety; 5-methylcytosine;
hyperproliferative disorder; cancer, neurological disorder;
aberrant nitric oxide signalling; inflammation; tumour formation;
cytostatic; neuroprotective; antiinflammatory; antimicrobial.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  New antisense compounds targeted to nucleic acid molecules encoding DEXRAS1, useful for treating diseases associated with expression of DEXRAS1, e.g. neurological disorder or hyperproliferative disorder such
                                                                                                                                  Gaps
                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                        Human DEXRAS1 DNA antisense oligonucleotide target region #24.
                                                                                             Length 20;
                                                                                                                                0; Indels
                                                       Sequence 20 BP; 13 A; 2 C; 1 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Sequence 20 BP; 4 A; 7 C; 4 G; 5 T; 0 U; 0 Other;
                                                                                           0.4%; Score 15; DB 1; L
100.0%; Pred. No. 2.4e+02;
iive 0; Mismatches 0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Example 15; SEQ ID NO 114; 62pp; English.
e.g., lung, brain, heart, kidney, etc, prevent any unwanted effects due to it
                                                                                                                                                                                                                                                                                                         踞
                                                                                                                                                                      3305 TTTTTATTTATAT 3319
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            28-JUN-2002; 2002US-00185035.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 28-JUN-2002; 2002US-00185035.
                                                                                                                                                                                                                                                                                                         ADI66983 standard; DNA; 20
                                                                                                                                                                                                            rrrrarrriarar
                                                                                       Query Match 0.4%
Best Local Similarity 100.0
Matches 15, Conservative
                                                                                                                                                                                                                                                                                                                                                                                  22-APR-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                WPI; 2004-081730/08.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Monia BP, Dobie KW
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  US2004005706-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Homo sapiens
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      08-JAN-2004.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 as cancer.
                                                                                                                                                                                                                                                                                                                                             ADI66983;
                                                                                                                                                                                                          15
                                                                                                                                                                                                                                                               SSXS
                                                                                                                                                                        ઠે
                                                                                                                                                                                                        셤
```

```
The invention relates to an antisense oligonucleotide targeted to a nucleic acid encoding the human DERRASI polypeptide which specifically hybridises with the nucleic acid molecule and inhibits the expression of DERRASI. The antisense oligonucleotide comprises at least one modified sugar molety, preferably a 2'-0-methoxyethyl sugar molety, or at least one modified sugar molety, preferably a 2'-0-methoxyethyl sugar molety, or at least one modified nucleobase comprising a 5-metholycytosine. The antisense oligonucleotides and compounds are useful for inhibiting the artisense oligonucleotides and compounds are useful for inhibiting the expression of DEXRASI e.g. hyperproliferative disorders such as cancer, neurological disorders or a disease or condition arising from aberrant nitric oxide signalling. The antisense compounds are also useful as research reagents and kits or in diagnostic, therapeutic and prophylactic applications. This sequence represents an antisense oligonucleotide of the invention. This sequence represents an antisense
                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ö
                                                                                                                                                                                                                                                                                                                                 Human, DEXRAS1; ss; antisense oligonucleotide; phosphorothioate linkage;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     New antisense compounds targeted to nucleic acid molecules encoding DEXRAS1, useful for treating diseases associated with expression of DEXRAS1, e.g. neurological disorder or hyperproliferative disorder such
                                Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Gaps
                                                                                                                                                                                                                                                                                                                                               2'-0-methoxyethyl sugar moiety; 5-methylcytosine; hyperproliferative disorder; cancer; neurological disorder; aberrant nitric oxide signalling; inflammation; tumour formation; cytostatic; neuroprotective; antinflammatory; antimicrobial.
                                ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ;
0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Length 20;
                                Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Indels
                                  ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Sequence 20 BP; 5 A; 4 C; 7 G; 4 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                0.4%; Score 15; DB 1; Le
.00.0%; Pred. No. 2.4e+02;
.ve 0; Mismatches 0;
              Pred. No. 2.4e+02; 
; Mismatches 0;
                                                                                                                                                                                                                                                                                              Human DEXRAS1 DNA antisense oligonucleotide #45.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Example 15; SEQ ID NO 57; 62pp; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 0.10,
100.0%; Pre-
10.001
14. $0.001
                                                                                                                                                                                         ADI66926 standard; DNA; 20 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    2509 AGGIGGAGCIGIACC 2523
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              28-JUN-2002; 2002US-00185035.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              28-JUN-2002; 2002US-00185035.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1 AGGIGGAGCIGIACC 15
                                                                                                   9
                                                                                                                                                                                                                                                            22-APR-2004 (first entry)
                              15; Conservative
                                                                 2509 AGGTGGAGCTGTACC
                                                                                                 20 AGGTGGAGCTGTACC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               (ISIS-) ISIS PHARM INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Dobie KW;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WPI; 2004-081730/08.
                Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Local Similarity
tes 15; Conserv
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         US2004005706-A1.
                                                                                                                                                                                                                                                                                                                                                                                                                                          Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           08-JAN-2004
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Monia BP,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         as cancer
                                                                                                                                                                                                                           ADI66926;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Query Match
                                Matches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Matches
                                                                                                                                                                        AD166926
                                                                   ઠ
                                                                                                 요
                                                                                                                                                                                                           ò
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     g
```

```
BX551013 Glossina morsitans morsitans adult infected gut Glossina morsitans clone Tsell6al2_qlc, mRNA sequence.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Glossina morsitans morsitans Glossina morsitans morsitans morsitans follossina morsitans morsitans Glossina morsitans morsitans morsitans morsitans morsitans morsitans morsitans morsitans. Hippoboscoidea; Glossinidae; Glossinia, Glossinia, Glossinia, M. Kerhornou, A., Berriman, M., Lehane, M.J., Aksoy, S., Gibson, W., Kerhornou, A., Berriman, M., Hamilton, J., Soares, M.B., Bonaldo, M.F., Lehane, S. and Hall, N. Adult midgut expressed sequence tags from the tsetse fly Glossina morsitans and expression analysis of putative immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ACCESSION: BX551013
                                                                                                                             February 16, 2005, 09:29:15; Search time 0.001 Seconds (without alignments) 144.818 Million cell updates/sec
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.
                                                                                                                                                                                                                                                                        1 acagggccactgctgctcac......tctcactgacaggagaacta 3811
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Description
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Contact: Hall N
Pathogen Sequencing Unit
The Sanger Institute The Wellcome Trust Genome Campus
Hinxton, Cambridge, CB10 18A, UK
Request for clones, please contact: Mike Lehane
GenCore version 5.1.6
Copyright (c) 1993 - 2005 Compugen Ltd.
                                                                                                                                                                                                                                                                                                                                                                                                                                    ~
                                                                                                                                                                                                                                                                                                                                                                                                                             Total number of hits satisfying chosen parameters:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ALIGNMENTS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  SUMMARIES
                                                                                         OM nucleic - nucleic search, using sw model
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Post-processing: Minimum Match 0%
Maximum Match 100%
Listing first 1 summaries
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     response genes
Genome Biol. 4 (10), R63 (2003)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            19 1 BX551013
                                                                                                                                                                                                                                                                                                                   IDENTITY_NUC
Gapop 10.0 , Gapext 0.5
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    H
                                                                                                                                                                                                                                                                                                                                                                                     1 segs, 19 residues
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       BX551013.1 GI:33374827
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query
Match Length DB
                                                                                                                                                                                                                         us-10-001-863-3
3811
                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Minimum DB seq length: 8 Maximum DB seq length: 80
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           rstdb:*
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 4.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       14519198
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  BX551013
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  22881942
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Score
                                                                                                                                                                                                                                                                                                                   Scoring table:
                                                                                                                                                                                                                              Title:
Perfect score:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 15.8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            RESULT 1
BX551013
LOCUS
DEFINITION
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     VERSION
KEYWORDS
SOURCE
ORGANISM
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Database :
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   JOURNAL
MEDLINE
PUBMED
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 REFERENCE
AUTHORS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ACCESSION
                                                                                                                                                                                                                                                                           Sequence:
                                                                                                                                                                                                                                                                                                                                                                                     Searched:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   TITLE
                                                                                                                                    Run on:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          COMMENT
```

Result No.

· sandada di

```
/note="country: Zimbabwe; EST from adult gut infected with T.brucei"
                                                                                                                                                                                                                                                                                                        ö
Prof. M.J.Lehane
School of Biological Sciences,
University of Wales,
Bangor LL57 2UW
Bangor LL57 2UW
Band cones with suffix qlc are reverse primer reads starting at 5'
end of the cDNA all plc reads are from
the 3' end.
                                                                                                    Gaps
                                                                                                                                                                                                                                                                                                        ö
                                                                                                                                                                                                                                                                            Length 19;
                                                                                                                                                                                                                                                                                                       2; Indels
                                                                                                                                                                                                                                                                             DB 1;
                                                                                                                                                                                                                                                                            ; Score 15.8; DE; Pred. No. 0; 0; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                  Search completed: February 16, 2005, 09:29:16 Job time : 0.001 secs
                                                                                              Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                  1974 GTTGAACGAATGGAATGTG 1992
                                                                                                                                                                                                                                                                                                                                                            1 Gricaacciaicraicic 19
                                                                                                                                                                                                                                                                            0.4%;
                                                                                                                                                                                                                                                                   Query Match
Best Local Similarity 89.59
Marches 17, Conservative
                                                                                                              source
                                                                                                FEATURES
                                                                                                                                                                                                                                                                                                                                   ò
                                                                                                                                                                                                                                                                                                                                                          g
```

THIS PAGE BLANK (USPTC)